

=> E NOMURA MASAHIRO/AU 25

E1	5	NOMURA MASAHIDE/AU
E2	77	NOMURA MASAHICO/AU
E3	138 -->	NOMURA MASAHIRO/AU
E4	1	NOMURA MASAHIRO S/AU
E5	4	NOMURA MASAHISA/AU
E6	1	NOMURA MASAHORI/AU
E7	1	NOMURA MASAICHIRO/AU
E8	1	NOMURA MASAITI/AU
E9	20	NOMURA MASAJI/AU
E10	2	NOMURA MASAKASTU/AU
E11	344	NOMURA MASAKATSU/AU
E12	38	NOMURA MASAKAZU/AU
E13	40	NOMURA MASAKI/AU
E14	7	NOMURA MASAKO/AU
E15	11	NOMURA MASAMI/AU
E16	2	NOMURA MASAMICHI/AU
E17	6	NOMURA MASANAO/AU
E18	1	NOMURA MASANARI/AU
E19	1	NOMURA MASANIRO/AU
E20	2	NOMURA MASANO/AU
E21	4	NOMURA MASANOBU/AU
E22	12	NOMURA MASANORI/AU
E23	143	NOMURA MASAO/AU
E24	2	NOMURA MASAOMI/AU
E25	1	NOMURA MASARO/AU

=> S (E2 OR E3 OR E4) AND (BENZYLTHIA?)

	77	"NOMURA MASAHICO"/AU
	138	"NOMURA MASAHIRO"/AU
	1	"NOMURA MASAHIRO S"/AU
	227	BENZYLTHIA?
L7	3	("NOMURA MASAHICO"/AU OR "NOMURA MASAHIRO"/AU OR "NOMURA MASAHIRO S"/AU) AND (BENZYLTHIA?)

=> S (E2 OR E3 OR E4) AND (LIPID?)

	77	"NOMURA MASAHICO"/AU
	138	"NOMURA MASAHIRO"/AU
	1	"NOMURA MASAHIRO S"/AU
	277666	LIPID?
L8	11	("NOMURA MASAHICO"/AU OR "NOMURA MASAHIRO"/AU OR "NOMURA MASAHIRO S"/AU) AND (LIPID?)

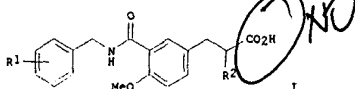
=> DIS L8 1 IBIB ABS

THE ESTIMATED COST FOR THIS REQUEST IS 2.29 U.S. DOLLARS
DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N:Y

L8 ANSWER 1 OF 11 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2002:428859 CAPLUS
 DOCUMENT NUMBER: 137:5998
 TITLE: Preparation of (phenylmethyl)alkanoic acid derivatives
 as PPAR.alpha. agonists for treatment of arteriosclerosis, obesity, diabetes, etc.
 INVENTOR(S): Miyachi, Hiroyuki; Nomura, Masahiro; Takahashi, Yukie; Tanase, Takahiro; Murakami, Kouji
 PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 52 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002044130	A1	20020606	WO 2001-JP10353	20011128
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BE, CA, CH, CN, CO, CR, CU, CY, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, T2, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, T2, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
PRIORITY APPLN. INFO.: JP 2000-363677 A			20001129	
OTHER SOURCE(S): MARPAT 137:5998				
GI				

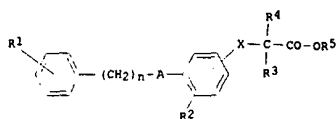
L8 ANSWER 1 OF 11 CAPLUS COPYRIGHT 2002 ACS (Continued)
 drugs (particularly in the liver), drugs preventing the progress of arteriosclerosis, anti-obesity drugs and remedies for diabetes.
 For example, 2-[[3-[[N-[(4-chlorophenyl)methyl]carbamoyl]-4-methoxyphenyl]methyl]butyric acid (II) was prepd. The PPAR.alpha. agonist activity of II was demonstrated.
 REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS
 RE FORMAT RECORD. ALL CITATIONS AVAILABLE IN THE



AB The title compds. I [R1 represents hydrogen, halogeno, hydroxy, 2-phenylethyl, 2-phenylethoxy, hydroxyphenoxy or benzyloxyphenoxy; and R2 represents lower (C1-4) alkyl] are prepd. I are lipid-lowering

L8 ANSWER 2 OF 11 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2002:428856 CAPLUS
 DOCUMENT NUMBER: 137:20225
 TITLE: Preparation of phenylmethylalkanoic acid
 derivatives
 treatment of as PPAR.alpha. agonists useful in the
 hyperlipidemia, arteriosclerosis, diabetes, and
 obesity
 INVENTOR(S): Miyachi, Hiroyuki; Momura, Masahiro;
 Murakami, Kouji
 PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 67 pp.
 CODEN: PIKXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE
 WO 2002044127 A1 20020606 WO 2001-JP10355 20011128
 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA,
 CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD,
 GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
 LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ,
 OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
 TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU,
 TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT,
 BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT,
 SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
 TD, TG
 PRIORITY APPLN. INFO.: JP 2000-363679 A 20001129
 OTHER SOURCE(S): MARPAT 137:20225
 GI



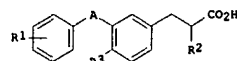
AB The title compds. I [R¹ represents trifluoromethyl, optionally substituted

L8 ANSWER 2 OF 11 CAPLUS COPYRIGHT 2002 ACS (Continued)
 phenoxy, etc.; R₂ represents hydrogen or lower alkoxy; R₃, R₄ and
 R₅ represent each hydrogen or lower alkyl; A represents NHCO or CONH;
 X is located at the para-position relative to A and represents oxygen or
 sulfur, or X is located at the para-position relative to R₂ and
 represents oxygen or sulfur; and n is an integer of from 0 to 2], useful as
 PPAR.alpha. agonists (no data) for the treatment of hyperlipidemia,
 arteriosclerosis, diabetes, and obesity, are prepd. For example,
 2-[[4-[[N-[[4-(trifluoromethyl)phenyl]methyl]carbamoyl]-3-
 methoxyphenyl]methyl]butyric acid was prepd.
 REFERENCE COUNT: 14 THERE ARE 14 CITED REFERENCES AVAILABLE
 FOR THIS
 RE FORMAT RECORD. ALL CITATIONS AVAILABLE IN THE

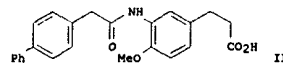
LB ANSWER 3 OF 11 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2001:886030 CAPLUS
 DOCUMENT NUMBER: 136:19941
 TITLE: Preparation of phenylpropionic acid
 derivatives as
 PPAR.alpha. activators effective as
 antiarteriosclerotics
 INVENTOR(S): Miyachi, Hiroyuki; Nomura, Masahiro;
 Takahashi, Yukie; Tanase, Takahiro; Murakami,
 Kouji;
 PATENT ASSIGNEE(S): Suzuki, Masahiro
 Kyorin Pharmaceutical Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 115 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001092201	A1	20011206	WO 2001-JP4385	20010525
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			JP 2000-158424 A	20000529
PRIORITY APPLN. INFO.: OTHER SOURCE(S):		MARPAT 136:19941		

LB ANSWER 3 OF 11 CAPLUS COPYRIGHT 2002 ACS (Continued)



I NO



AB Title compds. [I: R1 = alkyl, alkoxy, trifluoromethyl, trifluoromethoxy, Ph, phenoxy, benzyloxy; R2 = H, alkyl, alkoxy; R3 = alkoxy; A = CH2CONH, NHCOCH2, CH2CH2CO, CH2CH2CH2, CH2CH2O, CONHCH2, CH2NHCH2, COCH2O, OCH2CO, COCH2NH, NHCH2CO], stereoisomers, and pharmaceutically acceptable salts, which bind to human peroxisome proliferator activated receptor .alpha. (PPAR.alpha.) as ligand to activate the receptor and thereby exhibit a potent lipid-decreasing effect, are prepd. as antiarteriosclerotics. Thus, the title compd. II was prepd. and biol. tested for transcription activation effect with EC50(.mu.mol/L) = 0.05.
 REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L8 ANSWER 4 OF 11 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2001:511907 CAPLUS
 DOCUMENT NUMBER: 136:256870
 TITLE: Effects of idoxifene and estradiol on
 NF-.kappa.B activation in cultured rat hepatocytes
 undergoing oxidative stress
 AUTHOR(S): Omoya, Toshihiro; Shimizu, Ichiro; Zhou, Yajun;
 Guangming; Okamura, Yoshihito; Inoue, Hiroshi; Lu,
 Itonaga, Mina; Honda, Hirohito; Momura,
 Masahiro; Ito, Susumu
 CORPORATE SOURCE: Second Department of Internal Medicine,
 Tokushima University School of Medicine, Tokushima,
 770-8503, Japan
 SOURCE: Liver (Copenhagen, Denmark) (2001), 21(3),
 183-191
 PUBLISHER: Munksgaard International Publishers Ltd.
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Background/Aims: Idoxifene is a tissue-specific selective estrogen
 receptor modulator. Estradiol is a potent endogenous antioxidant,
 and nuclear factor .kappa.B (NF-.kappa.B) is a key transcription
 factor that induces multiple genes in response to inflammation or oxidative
 stress. The aim of this study was to explore the inhibitory effects of
 idoxifene and estradiol on NF-.kappa.B activation in hepatocytes in a state
 of oxidative stress. Methods: Lipid peroxidn. was induced in
 cultured rat hepatocytes by incubation with ferric
 nitrilotriacetate soln. NF-.kappa.B activity was evaluated by electrophoretic mobility
 shift assay. Results: The oxidative stress-induced activation of
 NF-.kappa.B and degrdn. of I.kappa.B-.alpha. were maximal at 3-5 h, with an
 increase in lactate dehydrogenase (LDH) and malondialdehyde (MDA) secretion
 into the culture medium. Treatment with idoxifene and estradiol
 inhibited I.kappa.B-.alpha. degrdn. and NF-.kappa.B activation through the
 attenuation of hepatocyte oxidative bursts and decreased
 extracellular levels of LDH and MDA. In addn., idoxifene and estradiol inhibited
 lipid peroxidn. in rat liver mitochondria. A potent NF-.kappa.B
 inhibitor, pyrrolidine dithiocarbamate, prevented NF-.kappa.B
 activation by inhibition of I.kappa.B-.alpha. degrdn. and decreased LDH and
 MDA

L8 ANSWER 4 OF 11 CAPLUS COPYRIGHT 2002 ACS (Continued)
 levels, suggesting that NF-.kappa.B might be a regulator in a
 genetic response to increase oxidative stress-induced hepatic injury.
 Conclusions: These findings suggest that idoxifene and estradiol
 function as antioxidants and protect hepatocytes from inflammatory cell
 injury.
 REFERENCE COUNT: 35 THERE ARE 35 CITED REFERENCES AVAILABLE
 FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE
 RE FORMAT

ND

L8 ANSWER 6 OF 11 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2001:152660 CAPLUS

DOCUMENT NUMBER: 134:193427

TITLE: Preparation of substituted

benzylthiazolidine-2,4-

peroxisome dione derivatives as agonists of human

proliferator-activated receptor

INVENTOR(S): Miyachi, Hiroyuki; Nomura, Masahiro; Tanase,

Takahiro; Murakami, Koji; Tsunoda, Masaki

PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan

SOURCE: PCT Int. Appl., 20 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001014351	A1	20010301	WO 2000-JP5521	20000818
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN,				
CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,				
ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LR, LS, LT, LU,				
LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG,				
SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM,				
AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE,				
CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,				
BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1207157 A1 20020522 EP 2000-953477 20000818				
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE,				
MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL				
PRIORITY APPLN. INFO.: JP 1999-235529 A 19990823				
JP 2000-242707 A 20000810				
WO 2000-JP5521 W 20000818				

OTHER SOURCE(S): MARPAT 134:193427

GI

L8 ANSWER 6 OF 11 CAPLUS COPYRIGHT 2002 ACS (Continued)

AB The title compds. represented by general formula (I; wherein A

represents

optionally substituted Ph, optionally substituted phenoxy or

optionally

substituted benzyloxy), pharmaceutically acceptable salts thereof

and

hydrates of the same are prepd. These compds. are capable of, as

a ligand of human peroxisome proliferator-activated receptor (PPAR),

enhancing the transcriptional activity of the receptor and showing effects of

lowering

blood sugar level and lowering lipid level. Thus,

5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxybenzoic acid, Et3N,

and

CH2Cl2 were mixed, treated with Et chlorocarbonate under

ice-cooling, and

stirred for 10 min under ice-cooling, followed by adding a soln. of

4-benzyloxybenzylamine in CH2Cl2, and the resulting mixt. was

stirred at

room temp. for 2 h to give 77%

N-[(4-benzyloxyphenyl)methyl]-5-[(2,4-

dioxothiazolidin-5-yl)methyl]-2-methoxybenzamide (II). II and I

(A = PhO)

enhanced the transcriptional activity of human PPAR.alpha. in CHO

cells

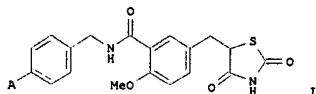
with EC50 of 0.44 and 0.24 .mu.M, resp.

REFERENCE COUNT: 27 THERE ARE 27 CITED REFERENCES AVAILABLE

FOR THIS

RE FORMAT

RECORD. ALL CITATIONS AVAILABLE IN THE



L8 ANSWER 7 OF 11 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2000:881114 CAPLUS

DOCUMENT NUMBER: 134:29211

TITLE: Preparation of
phenylmethylcarbamoylphenylpropionic
acid derivatives as human peroxisome
proliferator-activated receptor-.alpha.

(PPAR-.alpha.)

agonists
INVENTOR(S): Nomura, Masahiro; Takahashi, Yukie; Tanase,
Takahiro; Miyachi, Hiroyuki; Tsunoda, Masaki;

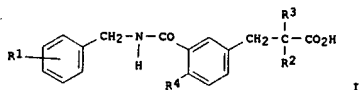
Tomohiro; Murakami, Koji
Ide, Kyorin Pharmaceutical Co., Ltd., Japan
PATENT ASSIGNEE(S): ECT Int. Appl., 90 pp.
SOURCE: CODEN: PIXXD2

DOCUMENT TYPE: Patent
LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000075103	A1	20001214	WO 2000-JP3707	20000608
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN,				
CR, CU,	CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,			
ID, IL,	IN, IS, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV,			
MD, MG,	MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI,			
SK, SL,	TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ,			
BY, KG,	KZ, MD, RU, TJ, TM			
CH, CY,	RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE,			
BF, BJ,	DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,			
	CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
JP 2001055367	A2	20010227	JP 2000-157600	20000529
BR 2000011734	A	20020305	BR 2000-11734	20000608
EP 1184366	A1	20020306	EP 2000-935582	20000608
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE,				
MC, PT,	IE, SI, LT, LV, FI, RO			
NO 2001006001	A	20020211	NO 2001-6001	20011207
PRIORITY APPLN. INFO.:			JP 1999-162235	A 19990609
			JP 2000-157600	A 20000529
			WO 2000-JP3707	W 20000608
OTHER SOURCE(S):	MARPAT 134:29211			
GI				

L8 ANSWER 7 OF 11 CAPLUS COPYRIGHT 2002 ACS (Continued)



AB The title compds. I [R1 = alkyl, etc.; R2 = alkyl, alkoxy, etc.;
when R2
is alkyl, 2,2,2-trifluoroethyl, R3 is H, alkyl; when R2 is alkoxy,
phenoxy, etc., R3 is H; R4 = alkoxy] are prepd. I activate the
PPAR-.alpha. receptors and lower blood lipid (cholesterol and
neutral lipid) levels. 2-Ethyl-3-[4-methoxy-3-[N-[[4-
(trifluoromethyl)phenyl]methyl]carbamoyl]phenyl]propionic acid at
30 mg/kg
gave 55% decrease in total cholesterol in rats.

REFERENCE COUNT: 32 THERE ARE 32 CITED REFERENCES AVAILABLE
FOR THIS

RE FORMAT

RECORD. ALL CITATIONS AVAILABLE IN THE

L8 ANSWER 8 OF 11 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 1997:116453 CAPLUS
DOCUMENT NUMBER: 126:157499
TITLE: Preparation of N-substituted
dioxothiazolidylbenzamide

INVENTOR(S): derivatives as blood sugar lowering agents
Maeda, Toshio; ~~Nomura, Masahiro~~; Awano,
Katsuya; Kinoshita, Susumu; Sato, Hiroya;

Murakami,

PATENT ASSIGNEE(S): Kojii Tsuyoda, Masaki
Kyorin Sanyaku Kk. Japan
SOURCE: Jpn. Kokai Tokkyo Koho, 11 pp.
CODE: J000AF

DOCUMENT TYPE: Patent
LANGUAGE: Japanese

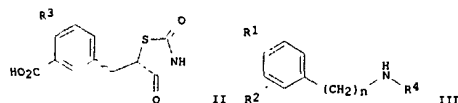
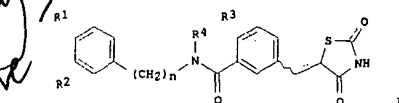
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 08333355	A2	19961217	JP 1995-159782	19950602

OTHER SOURCE(S): MARPAT 126:157499

GI



AB The title compds. (I; R1, R2 = H, C1-4 alkyl, C1-3 alkoxy, haloalkoxy, or haloalkyl, halo, OH, NO2, etc.; R3 = H, C1-3 alkoxy, halo, OH; R4 = H, C1-4 alkyl; dotted line = single or double bond; n = 0-2) are prepd. by reacting benzoic acid derivs. (II; R3, dotted line = same as above) with amines (III; R1, R2, R4, n = same as above). I, possessing blood sugar and lipid lowering activities, are useful for diabetes mellitus and hyperlipemia. Thus, 5-(2,4-dioxothiazolidyl-5-ylidene)methyl-2-

L8 ANSWER 8 OF 11 CAPLUS COPYRIGHT 2002 ACS (Continued)
methoxybenzoic acid was reacted with 4-tert-butylaniline in the presence of Et3N and NCP(O)(OEt)2 to give 99% I (R1 = 4-tert-BuC6H4, R3 = 2-MeO, R2 = R4 = H, dotted line = double bond, n = 0). I (R1 = R2 = 4-CF3, R3 = 6-MeO, R4 = Et, dotted line = single bond, n = 1) at 10 mg/kg showed 31% blood sugar lowering activity when tested on mice p.o. in vivo.

same assignee

103?

ACCESSION NUMBER: 1988:548377 CAPLUS
DOCUMENT NUMBER: 109:148377
TITLE: Effect of the dietary
.alpha.-linolenate/linoleate
balance on lipid compositions and learning
ability of rats. II. Discrimination process,
extinction process, and glycolipid compositions
Yamamoto, Nobuhiro; Hashimoto, Atsusi;
AUTHOR(S):
Takemoto, Yasuhiko; Okuyama, Harumi; Nomura, Masahiko;
Kitejima, Rie; Togashi, Takako; Tamai, Yoichi
CORPORATE SOURCE: Fac. Pharm. Sci., Nagoya City Univ., Nagoya,
467,
Japan
SOURCE: J. Lipid Res. (1988), 29(8), 1013-21
CODEN: JLPRAW; ISSN: 0022-2275
DOCUMENT TYPE: Journal
LANGUAGE: English
AB Donryu strain rats through 2 generations were fed semi-purified
diets
supplemented with safflower seed oil (rich in linoleic acid) or
perilla
seed oil (rich in .alpha.-linolenic acid) or fed a conventional
lab. chow
(normal control diet). The brightness-discrimination learning
ability was
highest in the perilla oil-fed group, followed by the normal
group, and
then by the safflower group, extending the earlier observation in a
different strain of rat that .alpha.-linolenic acid is a factor in
maintaining high learning ability (Yamamoto, N. et al., 1987).
After the
brightness-discrimination learning test was administered,
extinction of
learning was measured. The time required for extinction was
significantly
longer in the safflower group than in either the perilla group or
the
normal diet group. Thus, the dietary .alpha.-linolenate/linoleate
balance
affected both learning and the extinction of learning. The
glycolipids of
the cerebrum, cerebellum, and olfactory lobe were analyzed.
Although the
fatty acid compns. of the sulfatide and gangliosides were
significantly
different in the 3 parts of the brain, relatively little
difference was
obsd. in the fatty acids of glycolipids between the safflower
group and
the perilla group, suggesting that gross changes in brain
glycolipids are
not responsible for the differences in learning abilities between
these
dietary groups.

NP

L8 ANSWER 10 OF 11 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 1987:195209 CAPLUS
 DOCUMENT NUMBER: 106:195209
 TITLE: Effect of dietary .alpha.-linolenate/linoleate
 balance on brain lipid compositions and learning
 ability of rats
 AUTHOR(S): Yamamoto, Nobuhiro; Saitoh, Masaki; Moriuchi,
 Atsuko; Nomura, Masahiko; Okuyama, Harumi
 CORPORATE SOURCE: Fac. Pharm. Sci., Nagoya City Univ., Nagoya,
 467, Japan
 SOURCE: J. Lipid Res. (1987), 28(2), 144-51
 CODEN: JLPRAW ISSN: 0022-2275
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Spontaneously hypertensive rats (SHR) and normotensive control,
 Wistar/Kyoto (WKY) rats through two generations were fed a
 semipurified diet supplemented either with safflower oil (rich in linoleate
 [60-33-3]) or with perilla oil (rich in .alpha.-linolenate [463-40-1]). The
 cerebral lipid contents and phospholipid compns. did not differ
 between the two dietary groups of SHR rats. There were also no
 differences in the unsatd./satd. ratios of individual
 phospholipids or the proportions of plasmalogens. However, the proportions of n-3 and
 n-6 fatty acids were significantly different. Decreases in the
 proportions of docosahexaenoate [22:6 (n-3)] [6217-54-5] in
 phosphatidylethanolamine and phosphatidylserine in the safflower oil group were compensated for
 with increases in the proportions of docosatetraenoic acid [22:4 (n-6)] [28874-58-0] and docosapentaenoic acid [22:5 (n-6)] [25182-74-5]
 as compared with the perilla oil group. These differences in
 phospholipidacyl chains were much smaller than the difference in
 the proportions of linoleate and .alpha.-linolenate of the diets. In a
 brightness-discrimination learning test, the total no. of
 responses to the pos. and neg. stimuli were less in the groups fed perilla oil.
 However, the .alpha.-linolenate-deficient group took longer to decrease the
 frequency of R- responses and therefore longer to learn the
 discrimination. Consequently, the correct response ratios were
 higher in the perilla oil groups than in the safflower oil groups. Thus, the
 dietary .alpha.-linolenate/linoleate balance influenced the
 (n-3)/(n-6) balance of polyenoic fatty acids differently among brain
 phospholipids.
 These changes in fatty acid compn. were accompanied by changes in
 the

L8 ANSWER 10 OF 11 CAPLUS COPYRIGHT 2002 ACS (Continued)
 brightness-discrimination learning ability in SHR and WKY rats,
 with rats fed a diet enriched in .alpha.-linolenate being superior in the
 correct response ratio.

L8 ANSWER 11 OF 11 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 1986:114062 CAPLUS
DOCUMENT NUMBER: 105:114062
TITLE: Effects of dl-.alpha.-tocopherol on lipid
peroxide
AUTHOR(S): Ebisu, Hiroshi; Koide, Tadashi; Nomura,
Masahiko; Nagata, Yutaka
CORPORATE SOURCE: Res. Off., Fukuyukai Hosp., Aichi, Japan
SOURCE: Igaku to Seibutsugaku (1985), 111(6), 343-6
CODEN: IGSBAL; ISSN: 0019-1604
DOCUMENT TYPE: Journal
LANGUAGE: Japanese
AB In mice fed with a high-lipid diet, the concns. of lipid
peroxides in the heart of animals injected with
dl-.alpha.-tocopherol
[2074-53-5] (15.0 mg/100 g) were lower than those in mice without
dl-.alpha.-tocopherol.

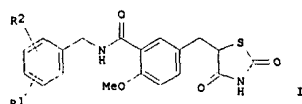
ND

Japan
37196
PCT

ICATION NO.

000-JP5522
R, BY, CA,
GE, GH, GM

RE. FORMAT

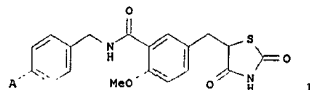


AB The title compds. (I), pharmaceutically acceptable salts thereof
and

L12 ANSWER 2 OF 4 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2001:152660 CAPLUS
 DOCUMENT NUMBER: 134:193427
 TITLE: Preparation of substituted benzylthiazolidine
 -2,4-dione derivatives as agonists of human
 peroxisome
 INVENTOR(S): No proliferator-activated receptor
 Takahiro: Miyachi, Hiroyuki; Nomura, Masahiro; Tanase,
 Murakami, Koji; Tsunoda, Masaki
 PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 20 pp.
 CODEN: PIXKD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001014351	A1	20010301	WO 2000-JP5521	20000818
W:	AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GR, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
EP 1207157	A1	20020522	EP 2000-953477	20000818
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL			
PRIORITY APPLN. INFO.:			JP 1999-235529 A 19990823	
			JP 2000-242707 A 20000810	
			WO 2000-JP5521 W 20000818	
OTHER SOURCE(S):		MARPAT 134:193427		
G1				

L12 ANSWER 2 OF 4 CAPLUS COPYRIGHT 2002 ACS (Continued)
 AB The title compds. represented by general formula (I), wherein A represents optionally substituted Ph, optionally substituted phenoxy or optionally substituted benzyl, pharmaceutically acceptable salts thereof and hydrates of the same are prep'd. These compds. are capable of, as a ligand of human peroxisome proliferator-activated receptor (PPAR), enhancing the transcriptional activity of the receptor and showing effects of lowering blood sugar level and lowering lipid level. Thus, 5-[(2,4-dioxothiazolidin-3-yl)methyl]-2-methoxybenzoic acid, Et3N, and CH2Cl2 were mixed, treated with Et chlorocarbonate under ice-cooling, and stirred for 10 min under ice-cooling, followed by adding a soln. of 4-benzoyloxybenzylamine in CH2Cl2, and the resulting mixt. was stirred at room temp. for 2 h to give 77% N-[(4-benzoyloxyphenyl)methyl]-5-[(2,4-dioxothiazolidin-3-yl)methyl]-2-methoxybenzamide (II). II and I (A = PhO) enhanced the transcriptional activity of human PPAR.alpha. in CHO cells with EC50 of 0.44 and 0.24 .mu.M, resp.
 REFERENCE COUNT: 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

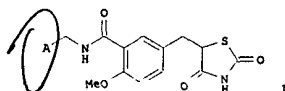


NO

L12 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2001:152659 CAPLUS
 DOCUMENT NUMBER: 134:178551
 TITLE: Preparation of substituted benzylthiazolidine
 -2,4-dione derivatives as ligands of human
 peroxisome
 proliferator-activated receptor
 INVENTOR(S): Fujimori, Shizuyoshi; Murakami, Koji;
 Tsunoda, Masaki
 PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 18 pp.
 CODEN: PIXX02
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001014350	A1	20010301	WO 2000-JP5520	20000818
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN,				
CR, CU,	CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,			
ID, IL,	IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,			
LV, MD,	MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG,			
SI, SK,	SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM,			
AZ, BY,	KG, KZ, MD, RU, TJ, TM			
CH, CY,	RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE,			
BF, BJ,	DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,			
	CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
EP 1207156	A1	20020522	EP 2000-953476	20000818
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE,				
MC, PT,	IE, SI, LT, LV, FI, RO, MK, CY, AL			
PRIORITY APPLN. INFO.:	JP 1999-235528	A	19990823	
	WO 2000-JP5520	W	20000818	

GI



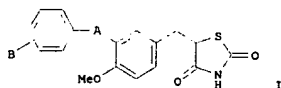
AB The title compds. (I; wherein A represents pyridyl or cyclohexyl),
 pharmaceutically acceptable salts thereof and hydrates of the same
 are

L12 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2002 ACS (Continued)
 prepd. These compds. are capable of, as a ligand of human
 peroxisome
 proliferator-activated receptor (PPAR), enhancing the
 transcriptional
 activity of the receptor and showing effects of lowering blood
 sugar level
 and lowering lipid level. Thus,
 5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-
 methoxybenzoic acid, 2-picolylamine, 1-ethyl-3-(3-
 dimethylaminopropyl)carbodiimide hydrochloride, and DMF were
 stirred at
 room temp. overnight to give 20% I (A = 2-pyridyl) (II). II and I
 (A =
 4-pyridyl) enhanced the transcriptional activity of human
 PPAR.alpha. in
 CHO cells with EC50 of 0.353 and 0.235 .mu.M, resp., and that of
 human
 PPAR.gamma. with EC50 of 0.30 and 0.14 .mu.M, resp.
 REFERENCE COUNT: 20 THERE ARE 20 CITED REFERENCES AVAILABLE
 FOR THIS
 REFORMAT RECORD. ALL CITATIONS AVAILABLE IN THE

NO - NO

L12 ANSWER 4 OF 4 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2001:152658 CAPLUS
 DOCUMENT NUMBER: 134:193426
 TITLE: Preparation of substituted benzylthiazolidine-2,4-dione derivatives as agonists for peroxisomal proliferator activated receptor (PPAR)
 INVENTOR(S): Miyachi, Hiroyuki; Nomura, Masahiro; Tanase, Takahiro;
 PATENT ASSIGNEE(S): Mirakami, Koji; Tsunoda, Masaki
 Kyorin Pharmaceutical Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 49 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001014349	A1	20010301	WO 2000-JP5519	20000818
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1213287	A1	20020612	EP 2000-953475	20000818
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL				
PRIORITY APPLN. INFO.: JP 1999-235527 A 19990823				
JP 2000-242706 A 20000810				
WO 2000-JP5519 W 20000818				
OTHER SOURCE(S): MARPAT 134:193426				
GI				



L12 ANSWER 4 OF 4 CAPLUS COPYRIGHT 2002 ACS (Continued)
 AB The title compds. (I) or pharmaceutically acceptable salts thereof, or hydrates of both (wherein A is CH₂CONH, NHCONH, CH₂CH₂CO, or NHCOCH₂; B is C1-4 lower alkyl, C1-3 lower alkoxy, halogeno, trifluoromethyl, trifluoromethoxy, substituted or unsubstituted Ph, substituted or unsubstituted phenoxy, or substituted or unsubstituted benzyloxy) are prepd. These compds. bind as ligands to human peroxisome proliferator-activated receptor (PPAR) to thereby activate the receptor and exert antihyperglycemic and antihyperlipidemic effects. Thus, 5-[(3-amino-4-methoxyphenyl)methyl]thiazolidine-2,4-dione was mixed with 5 mL THF, followed by adding 0.236 mL 4-trifluoromethyl isocyanate with stirring at room temp., and the stirring was continued at room temp. for 6 h, and left to stand overnight to give 57% 5-[[4-methoxy-3-[(4-(trifluoromethyl)phenyl)ureido]phenyl)methyl]thiazolidine-2,4-dione (II). II showed the transcription-activating activity for human PPAR.alpha. and PPAR.gamma. in CHO cells with EC₅₀ of 0.55 and 0.43 .mu.M, resp. REFERENCE COUNT: 36 THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> S (E2 OR E3 OR E4) AND (PPAR?)
172 "MURAKAMI KOICHI"/AU
239 "MURAKAMI KOJI"/AU
1 "MURAKAMI KOJIMA MASAYA"/AU
2767 PPAR?
L13 21 ("MURAKAMI KOICHI"/AU OR "MURAKAMI KOJI"/AU OR "MURAKAMI
KOJIMA
MASAYA"/AU) AND (PPAR?)

=> E TSUNODA MASAKI/AU 25
E1 2 TSUNODA MASAICHI/AU
E2 1 TSUNODA MASAKATSU/AU
E3 15 --> TSUNODA MASAKI/AU
E4 30 TSUNODA MASAKIYO/AU
E5 1 TSUNODA MASAKO/AU
E6 1 TSUNODA MASAKYO/AU
E7 16 TSUNODA MASAMI/AU
E8 2 TSUNODA MASANAO/AU
E9 1 TSUNODA MASANARI/AU
E10 13 TSUNODA MASAO/AU
E11 51 TSUNODA MASARU/AU
E12 13 TSUNODA MASASHI/AU
E13 6 TSUNODA MASATAKE/AU
E14 2 TSUNODA MASATO/AU
E15 1 TSUNODA MASATOSHI/AU
E16 2 TSUNODA MASAYA/AU
E17 1 TSUNODA MASAYOSHI/AU
E18 3 TSUNODA MASAYUKI/AU
E19 1 TSUNODA MASUKO/AU
E20 13 TSUNODA MAYUMI/AU
E21 1 TSUNODA MEGUMI/AU
E22 1 TSUNODA MICHIE/AU
E23 1 TSUNODA MICHIIHIKO/AU
E24 1 TSUNODA MIKI/AU
E25 7 TSUNODA MIKIO/AU

=> S (E3 OR E4) AND (PPAR?)
15 "TSUNODA MASAKI"/AU
30 "TSUNODA MASAKIYO"/AU
2767 PPAR?
L14 6 ("TSUNODA MASAKI"/AU OR "TSUNODA MASAKIYO"/AU) AND (PPAR?)

=> DIS L14 1 IBIB ABS
THE ESTIMATED COST FOR THIS REQUEST IS 2.29 U.S. DOLLARS
DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N:Y

L14 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:900080 CAPLUS
DOCUMENT NUMBER: 136:310816
TITLE: Design, synthesis and evaluation of substituted
phenylpropanoic acid derivatives as peroxisome
proliferator-activated receptor (PPAR)
activators: novel human PPAR
.alpha.-selective activators
AUTHOR(S): Miyachi, Hiroyuki; Nomura, Masahiro; Tanase,
Takahiro; Takehashi, Yukie; Ide, Tomohiro; Tsunoda,
Masaki; Murakami, Koji; Awano, Katsuya
CORPORATE SOURCE: Kyorin Pharmaceutical Co., Ltd., Discovery
Research Laboratories, Tochigi, Shimotsuga-gun,
Nogi-machi,
329-0114, Japan
SOURCE: Bioorganic & Medicinal Chemistry Letters
(2001),
Volume Date 2002, 12(1), 77-80
CODEN: BMCLE8; ISSN: 0960-894X
PUBLISHER: Elsevier Science Ltd.
DOCUMENT TYPE: Journal
LANGUAGE: English
AB A series of substituted phenylpropanoic acid derivs. was prepd. as
part of
a search for subtype-selective human peroxisome
proliferator-activated
receptor (PPAR) activators. Structure-activity relationship
studies indicated that the substituent at the .alpha.-position of
the
carboxyl group plays a key role in detg. the potency and the
selectivity
for PPAR transactivation.
REFERENCE COUNT: 19 THERE ARE 19 CITED REFERENCES AVAILABLE
FOR THIS
RECORD. ALL CITATIONS AVAILABLE IN THE
RE FORMAT

L14 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2001:152661 CAPLUS

DOCUMENT NUMBER: 134:193428

TITLE: Preparation of substituted

benzylthiazolidine-2,4-dione derivatives as agonists of human

peroxisome

proliferator-activated receptor
INVENTOR(S): Nomura, Masahiro; Murakami, Koji; Tsunoda,
Masaki; Takahashi, Yukie

PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan

SOURCE: PCT Int. Appl., 19 pp.

CODEN: PIXXD2

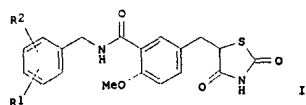
DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE
WO 2001014352 A1 20010301 WO 2000-JP5522 20000818
M: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN,
CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG,
SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM,
AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE,
CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,
BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
EP 1207158 A1 20020522 EP 2000-953478 20000818
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE,
MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL
PRIORITY APPLN. INFO.: JP 1999-235530 A 19990823
WO 2000-JP5522 W 20000818
OTHER SOURCE(S): MARPAT 134:193428
GI



L14 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2002 ACS (Continued)

AB The title compds. (I), pharmaceutically acceptable salts thereof
and

hydrates of the same (wherein R1 represents chloro, bromo, nitro,
trifluoromethoxy, ethoxy, propoxy or isopropoxy; and R2 represents
hydrogen or chloro) are prepd. These compds. are capable of, as a

ligand
of human peroxisome proliferator-activated receptor (PPAR),
enhancing the transcriptional activity of the receptor and showing

effects
of lowering blood sugar level and lowering lipid level; and a
process for

producing the same. Thus, 5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-
methoxybenzoic acid, Et3N, and CH2Cl2 were mixed, treated with Et
chlorocarbonate and stirred under ice-cooling for 10 min, treated

with
4-nitrobenzylamine, and then stirred at room temp. for 2 h to give

75%
N-[(4-nitrophenyl)methyl]-5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-
methoxybenzamide (II). II and I (R1 = 4-n-Pro, R2 = H) enhanced

the
transcriptional activity of human PPAR.alpha. in CHO cells with
EC50 of 0.53 and 0.11 .mu.M, resp.

REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE
FOR THIS

RE FORMAT RECORD. ALL CITATIONS AVAILABLE IN THE

L14 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2001:152660 CAPLUS
DOCUMENT NUMBER: 134:193427
TITLE: Preparation of substituted

benzylthiazolidine-2,4-dione derivatives as agonists of human

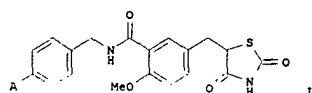
peroxisome proliferator-activated receptor
INVENTOR(S): Miyachi, Hiroyuki; Nomura, Masahiro; Tanase, Takahiro;

PATENT ASSIGNEE(S): Murakami, Koji; Tsunoda, Masaki
SOURCE: Kyorin Pharmaceutical Co., Ltd., Japan
PCT Int. Appl., 20 pp.

DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001014351	A1	20010301	WO 2000-JP5521	20000818
W:	AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU,			
	CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL,			
	IN, IS, JP, KE, KG, KP, KR, KZ, LC, LM, LR, LS, LT, LU, LV, MD,			
	MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK,			
	SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY,			
	KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,			
	DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,			
	CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
EP 1207157	A1	20020522	EP 2000-953477	20000818
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,			
	IE, SI, LT, LV, FI, RO, MK, CY, AL			
PRIORITY APPLN. INFO.:			JP 1999-235529	A 19990823
			JP 2000-242707	A 20000810
			WO 2000-JP5521	W 20000818
OTHER SOURCE(S):				
GI				



L14 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2002 ACS (Continued)

AB The title compds. represented by general formula (I; wherein A represents optionally substituted Ph, optionally substituted phenoxy or optionally substituted benzyloxy), pharmaceutically acceptable salts thereof and hydrates of the same are prepd. These compds. are capable of, as

a ligand of human peroxisome proliferator-activated receptor (PPAR), enhancing the transcriptional activity of the receptor and showing effects of lowering blood sugar level and lowering lipid level. Thus, 5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxybenzoic acid, Et3N,

and CH2Cl2 were mixed, treated with Et chlorocarbonate under ice-cooling, and stirred for 10 min under ice-cooling, followed by adding a soln. of 4-benzyloxybenzylamine in CH2Cl2, and the resulting mixt. was stirred at room temp. for 2 h to give 77%

N-[(4-benzyloxyphenyl)methyl]-5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxybenzamide (II). II and I (A = PhO)

enhanced the transcriptional activity of human PPAR.alpha. in CHO cells with EC50 of 0.44 and 0.24 .mu.M, resp.

REFERENCE COUNT: 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS

RE FORMAT RECORD. ALL CITATIONS AVAILABLE IN THE

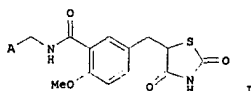
L14 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2001:152659 CAPLUS
DOCUMENT NUMBER: 134:178551
TITLE: Preparation of substituted
benzylthiazolidine-2,4-

peroxisome dione derivatives as ligands of human
proliferator-activated receptor
INVENTOR(S): Fujimori, Shizuyoshi; Murakami, Koji; Tsunoda,
Masaki
PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
SOURCE: PCT Int. Appl., 18 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001014350	A1	20010301	WO 2000-JP5520	20000818
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GM, GR, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LI, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1207156 A1 20020522 EP 2000-953476 20000818				
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL				
PRIORITY APPLN. INFO.: JP 1999-235528 A 19990823				
WO 2000-JP5520 W 20000818				

GI



AB The title compds. (I: wherein A represents pyridyl or cyclohexyl),
pharmaceutically acceptable salts thereof and hydrates of the same
are

L14 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2002 ACS (Continued)

prepd. These compds. are capable of, as a ligand of human
peroxisome
proliferator-activated receptor (PPAR), enhancing the
transcriptional activity of the receptor and showing effects of
lowering
blood sugar level and lowering lipid level. Thus, 5-[(2,4-
dioxothiazolidin-5-yl)methyl]-2-methoxybenzoic acid,
2-picolyamine,
1-ethyl-3-(3-dimethylaminopropyl)carbodiimide hydrochloride, and
DMF were
stirred at room temp. overnight to give 20% I (A = 2-pyridyl)
(II). II
and I (A = 4-pyridyl) enhanced the transcriptional activity of
human
PPAR.alpha. in CHO cells with EC50 of 0.353 and 0.235 .mu.M,
resp., and that of human PPAR.gamma. with EC50 of 0.30 and 0.14
.mu.M, resp.

REFERENCE COUNT: 20 THERE ARE 20 CITED REFERENCES AVAILABLE
FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE
RE FORMAT

L14 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2002 ACS

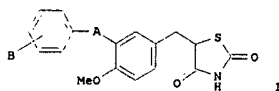
ACCESSION NUMBER: 2001:152658 CAPLUS
DOCUMENT NUMBER: 134:193426
TITLE: Preparation of substituted

benzylthiazolidine-2,4-dione derivatives as agonists for peroxisomal proliferator activated receptor (PPAR)
Miyachi, Hiroyuki; Nomura, Masahiro; Tanase, Takahiro;

INVENTOR(S): Murakami, Koji; Tsunoda, Masaki
PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
SOURCE: PCT Int. Appl., 49 pp.

DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001014349	A1	20010301	WO 2000-JP5519	20000818
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GR, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1213287	A1	20020612	EP 2000-953475	20000818
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL				
PRIORITY APPLN. INFO.: JP 1999-235527 A 19990823 JP 2000-242706 A 20000810 WO 2000-JP5519 W 20000818				
OTHER SOURCE(S): MARPAT 134:193426 GI				



L14 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2002 ACS (Continued)
AB The title compds. (I) or pharmaceutically acceptable salts thereof, or hydrates of both (wherein A is CH₂CONH, NHCONH, CH₂CH₂CO, or NHCOCH₂; B is C1-4 lower alkyl, C1-3 lower alkoxy, halogeno, trifluoromethyl, trifluoromethoxy, substituted or unsubstituted Ph, substituted or unsubstituted phenoxy, or substituted or unsubstituted benzyloxy) are

are prepd. These compds. bind as ligands to human peroxisome proliferator-activated receptor (PPAR) to thereby activate the receptor and exert antihyperglycemic and antihyperlipidemic effects.

Thus, 378 mg 5-[(3-amino-4-methoxyphenyl)methyl]thiazolidine-2,4-dione was mixed with 5 mL THF, followed by adding 0.236 mL 4-trifluoromethyl isocyanate with stirring at room temp., and the stirring was continued at room temp. for 6 h, and left to stand overnight to give 57%

5-[[4-methoxy-3-[(3-[4-(trifluoromethyl)phenyl]ureido)phenyl]methyl]thiazolidine-2,4-dione (II). II showed the transcription-activating activity for

human PPAR.alpha. and PPAR.gamma. in CHO cells with EC₅₀ of 0.55 and 0.43 .mu.M, resp.

REFERENCE COUNT: 36 THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS

RE FORMAT RECORD. ALL CITATIONS AVAILABLE IN THE

L14 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 2000:881114 CAPLUS

DOCUMENT NUMBER: 134:29211

TITLE: Preparation of

phenylmethylcarbamoylphenylpropionic acid derivatives as human peroxisome proliferator-activated receptor-.alpha. (PPAR-.alpha.) agonists

INVENTOR(S): Miyachi, Hiroyuki; Tsunoda, Masaki; Ide, Tomohiro; Murakami, Koji

PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan

SOURCE: PCT Int. Appl., 90 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

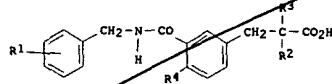
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

12-14-2000

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000075103	A1	20001214	WO 2000-JP3707	20000608
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CE, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
JP 2001055367	A2	20010227	JP 2000-157600	20000529
BR 2000011734	A	20020305	BR 2000-11734	20000608
EP 1184366	A1	20020306	EP 2000-935582	20000608
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
NO 2001006001	A	20020211	NO 2001-6001	20011207
PRIORITY APPLN. INFO.: JP 1999-162235 A 19990609				
JP 2000-157600 A 20000529				
WO 2000-JP3707 W 20000608				
OTHER SOURCE(S): MARPAT 134:29211				
GI				

L14 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2002 ACS (Continued)



AB The title compds. I (R1 = alkyl, etc.; R2 = alkyl, alkoxy, etc.; when R2 is alkyl, 2,2,2-trifluoroethyl, R3 is H, alkyl; when R2 is alkoxy, phenoxy, etc., R3 is H; R4 = alkoxy) are prep. I activate the PPAR-.alpha. receptors and lower blood lipid (cholesterol and neutral lipid) levels. 2-Ethyl-3-[4-methoxy-3-[N-[[4-(trifluoromethyl)phenyl]methyl]carbamoyl]phenyl]propionic acid at 30 mg/kg gave 55% decrease in total cholesterol in rats.

REFERENCE COUNT: 32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> E TAKAHASHI YUKIE/AU 25

E1	30	TAKAHASHI YUKI/AU
E2	1	TAKAHASHI YUKIBUMI/AU
E3	38	--> TAKAHASHI YUKIE/AU
E4	2	TAKAHASHI YUKIHARU/AU
E5	11	TAKAHASHI YUKIHIKO/AU
E6	92	TAKAHASHI YUKIHIRO/AU
E7	22	TAKAHASHI YUKIHISA/AU
E8	26	TAKAHASHI YUKIKO/AU
E9	26	TAKAHASHI YUKIMI/AU
E10	12	TAKAHASHI YUKINOBU/AU
E11	41	TAKAHASHI YUKINORI/AU
E12	324	TAKAHASHI YUKIO/AU
E13	2	TAKAHASHI YUKISHIGE/AU
E14	1	TAKAHASHI YUKISUKE/AU
E15	1	TAKAHASHI YUKITATSU/AU
E16	1	TAKAHASHI YUKITOMO/AU
E17	13	TAKAHASHI YUKITOSHI/AU
E18	31	TAKAHASHI YUKITSUGU/AU
E19	1	TAKAHASHI YUKIYO/AU
E20	1	TAKAHASHI YUKIYOSHI/AU
E21	60	TAKAHASHI YUKO/AU
E22	14	TAKAHASHI YUKOH/AU
E23	15	TAKAHASHI YUMI/AU
E24	1	TAKAHASHI YUMIE/AU
E25	1	TAKAHASHI YUMIHIRO/AU

=> S (E1 OR E2 OR E3) AND (PPAR?)

30	"TAKAHASHI YUKI"/AU
1	"TAKAHASHI YUKIBUMI"/AU
38	"TAKAHASHI YUKIE"/AU
2767	PPAR?

L15 6 ("TAKAHASHI YUKI"/AU OR "TAKAHASHI YUKIBUMI"/AU OR "TAKAHASHI YUKIE"/AU) AND (PPAR?)

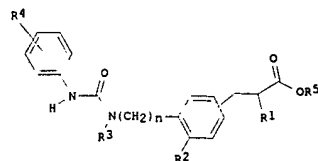
=> DIS L15 1 IBIB ABS

THE ESTIMATED COST FOR THIS REQUEST IS 2.29 U.S. DOLLARS

DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N:Y

L15 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2002:449639 CAPLUS
 DOCUMENT NUMBER: 137:33138
 TITLE: Preparation of ureidophenylalkanoic acid and ureidoalkylphenylalkanoic acid derivatives as human peroxisome proliferator-activated receptor .alpha. (PPAR.alpha.) agonists
 INVENTOR(S): Miyachi, Hiroyuki; Takahashi, Yukie; Murakami, Kouji
 PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 35 pp. CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002046146	A1	20020613	WO 2001-JP10563	20011204
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
PRIORITY APPLN. INFO.: JP 2000-369371 A 20001205				
OTHER SOURCE(S): MARPAT 137:33138				
GI				



NO

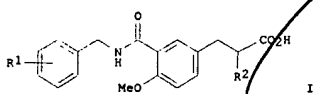
I

L15 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2002 ACS (Continued)

AB The title compds. I [R1, R3 and R5 represent each hydrogen or lower alkyl; R2 represents hydrogen or lower alkoxy; R4 represents hydrogen, trifluoromethyl, lower alkoxy, halogeno, optionally substituted phenoxy or benzyloxy; n is an integer of from 0 to 3; and the carboxylate substituent is located at the para-position relative to R2 or at the para-position relative to (CH2)n], useful as PPAR.alpha. agonists (no data), are prepd. I are useful in the treatment of diabetes, hyperlipidemia, obesity, and arteriosclerosis (no data). For example, 2-[[3-[3-[4-(trifluoromethyl)phenyl]ureido]-4-methoxyphenyl]methyl]butyric acid was prepd.
 REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L15 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2002:428859 CAPLUS
 DOCUMENT NUMBER: 137:5998
 TITLE: Preparation of (phenylmethyl)alkanoic acid derivatives
 as PPAR.alpha. agonists for treatment of arteriosclerosis, obesity, diabetes, etc.
 INVENTOR(S): Miyachi, Hiroyuki; Nomura, Masahiro; Takahashi, Yukio; Tanase, Takahiro; Murakami, Kouji
 PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 52 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002044130	A1	20020606	WO 2001-JP10353	20011128
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
PRIORITY APPLN. INFO.: MARPAT 137:5998			JP 2000-363677 A 20001129	
OTHER SOURCE(S):				
GI				



AB The title compds. I (R1 represents hydrogen, halogeno, hydroxy, 2-phenylethyl, 2-phenylethoxy, hydroxyphenoxy or benzyloxyphenoxy; and R2 represents lower (C1-4) alkyl) are prepd. I are lipid-lowering drugs

L15 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2002 ACS (continued)
 (particularly in the liver), drugs preventing the progress of arteriosclerosis, anti-obesity drugs and remedies for diabetes.
 For example, 2-[[3-[[N-[(4-chlorophenyl)methyl]carbamoyl]-4-methoxyphenyl]methyl]butyric acid (II) was prepd. The PPAR.alpha. agonist activity of II was demonstrated.
 REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS
 RE FORMAT RECORD. ALL CITATIONS AVAILABLE IN THE

L15 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:900080 CAPLUS
DOCUMENT NUMBER: 136:318816
TITLE: Design, synthesis and evaluation of substituted
phenylpropanoic acid derivatives as peroxisome
proliferator-activated receptor (PPAR)
activators: novel human PPAR
.alpha.-selective activators
AUTHOR(S): Miyachi, Hiroyuki; Nomura, Masahiro; Tanase,
Takahiro;
Takahashi, Tokie; Ide, Tomohiro; Tsunoda,
Masaki; Murakami, Koji; Awano, Katsuya
CORPORATE SOURCE: Kyorin Pharmaceutical Co., Ltd., Discovery
Research
Laboratories, Tochigi, Shimotsuga-gun,
Nogi-machi,
329-0114, Japan
SOURCE: Bioorganic & Medicinal Chemistry Letters
(2001),
Volume Date 2002, 12(1), 77-80
CODEN: BMCLES; ISSN: 0960-894X
PUBLISHER: Elsevier Science Ltd.
DOCUMENT TYPE: Journal
LANGUAGE: English
AB A series of substituted phenylpropanoic acid deriva. was prepd. as
part of
a search for subtype-selective human peroxisome
proliferator-activated
receptor (PPAR) activators. Structure-activity relationship
studies indicated that the substituent at the .alpha.-position of
the
carboxyl group plays a key role in detg. the potency and the
selectivity
for PPAR transactivation.
REFERENCE COUNT: 19 THERE ARE 19 CITED REFERENCES AVAILABLE
FOR THIS
RECORD. ALL CITATIONS AVAILABLE IN THE
RE FORMAT

ND

L15 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:886030 CAPLUS
DOCUMENT NUMBER: 136:19941
TITLE: Preparation of phenylpropionic acid
derivatives as

INVENTOR(S): **FFAR.alpha. activators effective as antiarteriosclerotics**
Miyachi, Hiroyuki; Nomura, Masahiro; Takahashi, Yukie; Tanase, Takahiro; Murakami, Kouji; Suzuki,

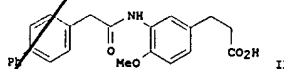
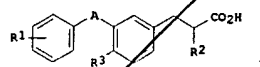
PATENT ASSIGNEE(S): Masahiro
SOURCE: Kyorin Pharmaceutical Co., Ltd., Japan
PCT Int. Appl., 115 pp.

DOCUMENT TYPE: CODEN: PIXXD2
LANGUAGE: Patent
FAMILY ACC. NUM. COUNT: Japanese
PATENT INFORMATION: 1

12.06.01

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001092201	A1	20011206	WO 2001-JP4385	20010525
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
PRIORITY APPLN. INFO.:			JP 2000-158424	A 20000529
OTHER SOURCE(S):			MARPAT 136:19941	
GI				

L15 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2002 ACS (Continued)

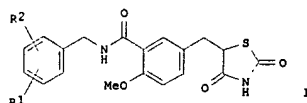


AB Title compds. [I: R1 = alkyl, alkoxy, trifluoromethyl, trifluoromethoxy, Ph, phenoxy, benzyloxy; R2 = H, alkyl, alkoxy; R3 = alkoxy; A = CH2CONH, NHCOCH2, CH2CH2CO, CH2CH2CH2, CH2CH2O, CONHCH2, CH2NHCH2, COCH2O, OCH2CO, COCH2NH, NHCH2CO], stereoisomers, and pharmaceutically acceptable salts, which bind to human peroxisome proliferator activated receptor .alpha. (FFAR.alpha.) as ligand to activate the receptor and thereby exhibit a potent lipid-decreasing effect, are prepd. as antiarteriosclerotics. Thus, the title compd. II was prepd. and biol. tested for transcription activation effect with EC50(.mu.mol/L) = 0.05.
REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS
RE FORMAT RECORD. ALL CITATIONS AVAILABLE IN THE

L15 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:152661 CAPLUS
DOCUMENT NUMBER: 134:193428
TITLE: Preparation of substituted
benzylthiazolidine-2,4-

peroxisome dione derivatives as agonists of human
proliferator-activated receptor
INVENTOR(S): Nomura, Masahiro; Murakami, Koji; Tsunoda,
Masaki; Takahashi, Tokie
PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
SOURCE: PCT Int. Appl., 19 pp.
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO DATE
WO 2001014352 A1 20010301 WO 2000JP5572 20000818
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, CA, CH, CN,
CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG,
SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM,
AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE,
CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,
BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
EP 1207158 A1 20020522 EP 2000-953478 20000818
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE,
MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL
PRIORITY APPLN. INFO.: JP 1999-235530 A 19990823
WO 2000-JP5522 W 20000818
OTHER SOURCE(S): MARPAT 134:193428
GI



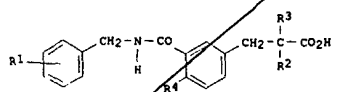
L15 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2002 ACS (Continued)

AB The title compds. (I), pharmaceutically acceptable salts thereof
and hydrates of the same (wherein R1 represents chloro, bromo, nitro,
trifluoromethoxy, ethoxy, propoxy or isopropoxy; and R2 represents
hydrogen or chloro) are prepd. These compds. are capable of, as a
ligand of human peroxisome proliferator-activated receptor (PPAR),
enhancing the transcriptional activity of the receptor and showing
effects of lowering blood sugar level and lowering lipid level; and a
process for producing the same. Thus, 5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-
methoxybenzoic acid, Et3N, and CH2Cl2 were mixed, treated with Et
chlorocarbonate and stirred under ice-cooling for 10 min, treated
with 4-nitrobenzylamine, and then stirred at room temp. for 2 h to give
75% N-[(4-nitrophenyl)methyl]-5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-
methoxybenzamide (II). II and I (R1 = 4-n-Pro, R2 = H) enhanced
the transcriptional activity of human PPAR.alpha. in CHO cells with
EC50 of 0.53 and 0.11 .mu.M, resp.
REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE
FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE
RE FORMAT

ACCESSION NUMBER: 2000:881114 CAPLUS
DOCUMENT NUMBER: 134:29211
TITLE: Preparation of
phenylmethylcarbamoylphenylpropionic
acid derivatives as human peroxisome
proliferator-activated receptor-.alpha. (PPAR
-.alpha.) agonists
INVENTOR(S): Nomura, Masahiro; Takahashi, Yukie; Tanase,
Takahiro; Miyachi, Hiroyuki; Tsunoda, Masaki;
Ide,
PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
SOURCE: PCT Int. Appl., 90 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000075103	A1	20001214	WO 2000-JP3707	20000608
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN,				
CR, CU,	CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,			
ID, IL,	IN, IS, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV,			
MD, MG,	MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI,			
SK, SL,	TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ,			
BY, KG,	KZ, MD, RU, TJ, TM			
CH, CY,	RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE,			
BF, BJ,	DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,			
	CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
JP 2001055367	A2	20010227	JP 2000-157600	20000529
BR 2000011734	A	20020305	BR 2000-11734	20000608
EP 1184366	A1	20020306	EP 2000-935582	20000608
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE,				
MC, PT,	IE, SI, LT, LV, FI, RO			
NO 2001006001	A	20020211	NO 2001-6001	20011207
PRIORITY APPLN. INFO.:			JP 1999-162235	A 19990609
			JP 2000-157600	A 20000529
			WO 2000-JP3707	W 20000608

OTHER SOURCE(S): MARPAT 134:29211
GI



AB The title compds. I [R1 = alkyl, etc.; R2 = alkyl, alkoxy, etc.;
when R2
is alkyl, 2,2,2-trifluoroethyl, R3 is H, alkyl; when R2 is alkoxy,
phenoxy, etc., R3 is H; R4 = alkoxy] are prepd. I activate the
PPAR-.alpha. receptors and lower blood lipid (cholesterol and
neutral lipid) levels. 2-Ethyl-3-[4-methoxy-3-[N-[[4-
(trifluoromethyl)phenyl)methyl]carbamoyl]phenyl]propionic acid at
30 mg/kg
gave 55% decrease in total cholesterol in rats.
REFERENCE COUNT: 32 THERE ARE 32 CITED REFERENCES AVAILABLE
FOR THIS
RE FORMAT
RECORD. ALL CITATIONS AVAILABLE IN THE

=> fil caplus		
COST IN U.S. DOLLARS	SINCE FILE	TOTAL
	ENTRY	SESSION
FULL ESTIMATED COST	267.97	414.57
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE	TOTAL
	ENTRY	SESSION
CA SUBSCRIBER PRICE	-36.55	-36.55

FILE 'CAPLUS' ENTERED AT 10:15:15 ON 12 AUG 2002
 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
 PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
 COPYRIGHT (C) 2002 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 12 Aug 2002 VOL 137 ISS 7
 FILE LAST UPDATED: 11 Aug 2002 (20020811/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

CAS roles have been modified effective December 16, 2001. Please check your SDI profiles to see if they need to be revised. For information on CAS roles, enter HELP ROLES at an arrow prompt or use the CAS Roles thesaurus (/RL field) in this file.

=> s PPAR?
 L16 2767 PPAR?

=> s bezylthia?
 L17 0 BEZYLTHIA?

=> s benzylthia?
 L18 227 BENZYLTHIA?

=> s l16 and l18 ful
 L19 5 L16 AND L18

=> d his

(FILE 'HOME' ENTERED AT 09:59:58 ON 12 AUG 2002)

FILE 'REGISTRY' ENTERED AT 10:01:05 ON 12 AUG 2002
 L1 STRUCTURE UPLOADED
 L2 46 S L1 FUL

L3 46 S L2 AND CAPLUS/LC

FILE 'CAPLUS' ENTERED AT 10:04:24 ON 12 AUG 2002

L4 32 S L2 FUL

L5 990331 S BLOOD?

L6 12 S L4 AND L5

E NOMURA MASAHIRO/AU 25

L7 3 S (E2 OR E3 OR E4) AND (BENZYLTHIA?)

L8 11 S (E2 OR E3 OR E4) AND (LIPID?)

E MURAKAMI KOJI/AU 25

L9 17 S (E2 OR E3 OR E4) AND (LIPID?)

L10 0 S (E2 OR E3 OR E4) AND (BEZYLTHI?)

L11 0 S (E2 OR E3 OR E4) AND (BEZYLTHIAZ?)

L12 4 S (E2 OR E3 OR E4) AND (BENZYLTHI?)

L13 21 S (E2 OR E3 OR E4) AND (PPAR?)

E TSUNODA MASAKI/AU 25

L14 6 S (E3 OR E4) AND (PPAR?)

E TAKAHASHI YUKIE/AU 25

L15 6 S (E1 OR E2 OR E3) AND (PPAR?)

FILE 'CAPLUS' ENTERED AT 10:15:15 ON 12 AUG 2002

L16 2767 S PPAR?

L17 0 S BEZYLTHIA?

L18 227 S BENZYLTHIA?

L19 5 S L16 AND L18 FUL

=> s l19 not l4

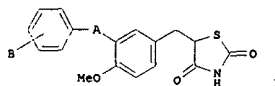
L20 2 L19 NOT L4

=> d l20 1-2 ibib abs hitstr

L20 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2001:653768 CAPLUS
 DOCUMENT NUMBER: 136:47867
 TITLE: Interaction between peroxisome proliferator-activated receptor .gamma. and its agonists: docking study of oximes having 5-benzyl-2,4-thiazolidinedione Iwata, Y.; Miyamoto, S.; Takamura, M.; Kasaaya, A.
 CORPORATE SOURCE: Exploratory Chemistry Research Laboratories, Sankyo Co., Ltd., Tokyo, Japan
 SOURCE: Journal of Molecular Graphics & Modelling 19(6), 536-542
 CODEN: JMGHFI; ISSN: 1093-3263
 PUBLISHER: Elsevier Science Inc.
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB The mol. modeling of oximes having 5-benzyl-2,4-thiazolidinedione moieties, agonists of the peroxisome proliferator-activated receptor .gamma. (PPAR.gamma.), was performed with respect to their structures complexed with the ligand binding domain of PPAR .gamma.. For each ligand mol., the 5-benzyl-2,4-thiazolidinedione head group was used as an anchor and the conformation of the rest of the mol. was searched for the most energetically favorable interaction with the receptor by systematic conformation search and manual modeling. Although both tail-up and tail-down configurations, which have been obsd. in the crystal structure of eicosapentaenoic acid when complexed with PPAR.gamma., appeared among the lowest energy structures for most of the compds., potent agonists were found to adopt a configuration similar to that of rosiglitazone when bound to PPAR.gamma., according to the crystal structure. The structure-activity relationships were analyzed based on the receptor-ligand interaction. The alkyl group and the arom. ring of the tail group of the ligands had hydrophobic interactions with the receptor, and these interactions were essential for the strong activity.
 REFERENCE COUNT: 20 THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS
 RE RECORD. ALL CITATIONS AVAILABLE IN THE
 RE FORMAT

L20 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2001:152658 CAPLUS
 DOCUMENT NUMBER: 134:193426
 TITLE: Preparation of substituted benzylthiazolidine -2,4-dione derivatives as agonists for peroxisomal proliferator activated receptor (PPAR)
 INVENTOR(S): Miyachi, Hiroyuki; Nomura, Masahiro; Tanase, Takahiro;
 PATENT ASSIGNEE(S): Murakami, Koji; Tsunoda, Masaki
 SOURCE: Kyorin Pharmaceutical Co., Ltd., Japan
 PCT Int. Appl., 49 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001014349	A1	20010301	WO 2000-JP5519	20000818
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, T2, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, GI, CH, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1213287	A1	20020612	EP 2000-953475	20000818
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL				
PRIORITY APPLN. INFO.:			JP 1999-235527	A 19990823
			JP 2000-242706	A 20000810
			WO 2000-JP5519	W 20000818
OTHER SOURCE(S):			MARPAT 134:193426	
GI				



L20 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2002 ACS (Continued)
 AB The title compds. (I) or pharmaceutically acceptable salts thereof, or hydrates of both (wherein A is CH2CONH, NHCONH, CH2CH2CO, or NHC(=O)CH2; B is C1-4 lower alkyl, C1-3 lower alkoxy, halogeno, trifluoromethyl, trifluoromethoxy, substituted or unsubstituted Ph, substituted or unsubstituted phenoxy, or substituted or unsubstituted benzylloxy)
 are prepd. These compds. bind as ligands to human peroxisome proliferator-activated receptor (PPAR) to thereby activate the receptor and exert antihyperglycemic and antihyperlipidemic effects.
 Thus, 378 mg 5-[(3-amino-4-methoxyphenyl)methyl]thiazolidine-2,4-dione was mixed with 5 mL THF, followed by adding 0.236 mL 4-trifluoromethyl isocyanate with stirring at room temp., and the stirring was continued at room temp. for 6 h, and left to stand overnight to give 57%
 5-[(4-methoxy-3-[(4-(trifluoromethyl)phenyl)ureido]phenyl)methyl]thiazolidine-2,4-dione (II). II showed the transcription-activating activity for human PPAR.alpha. and PPAR.gamma. in CHO cells with EC50 of 0.55 and 0.43 .mu.M, resp.
 REFERENCE COUNT: 36 THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS
 RE RECORD. ALL CITATIONS AVAILABLE IN THE
 RE FORMAT

foreign
priority
document

=> s dioxothia?
L21 247 DIOXOTHIA?

=> d his

(FILE 'HOME' ENTERED AT 09:59:58 ON 12 AUG 2002)

FILE 'REGISTRY' ENTERED AT 10:01:05 ON 12 AUG 2002

L1 STRUCTURE UPLOADED
L2 46 S L1 FUL
L3 46 S L2 AND CAPLUS/LC

FILE 'CAPLUS' ENTERED AT 10:04:24 ON 12 AUG 2002

L4 32 S L2 FUL
L5 990331 S BLOOD?
L6 12 S L4 AND L5
E NOMURA MASAHIRO/AU 25
L7 3 S (E2 OR E3 OR E4) AND (BENZYLTHIA?)
L8 11 S (E2 OR E3 OR E4) AND (LIPID?)
E MURAKAMI KOJI/AU 25
L9 17 S (E2 OR E3 OR E4) AND (LIPID?)
L10 0 S (E2 OR E3 OR E4) AND (BEZYLTHI?)
L11 0 S (E2 OR E3 OR E4) AND (BEZYLTHIAZ?)
L12 4 S (E2 OR E3 OR E4) AND (BENZYLTHI?)
L13 21 S (E2 OR E3 OR E4) AND (PPAR?)
E TSUNODA MASAKI/AU 25
L14 6 S (E3 OR E4) AND (PPAR?)
E TAKAHASHI YUKIE/AU 25
L15 6 S (E1 OR E2 OR E3) AND (PPAR?)

FILE 'CAPLUS' ENTERED AT 10:15:15 ON 12 AUG 2002

L16 2767 S PPAR?
L17 0 S BEZYLTHIA?
L18 227 S BENZYLTHIA?
L19 5 S L16 AND L18 FUL
L20 2 S L19 NOT L4
L21 247 S DIOXOTHIA?

=> s l21 and l16
L22 9 L21 AND L16

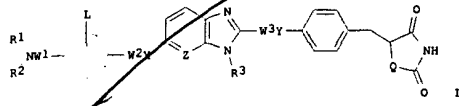
=> s l22 not l20
L23 8 L22 NOT L20

=> s l23 not l19
L24 5 L23 NOT L19

=> d l24 1-5 ibib abs hitstr

L24 ANSWER 1 OF 5 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2002:479961 CAPLUS
 DOCUMENT NUMBER: 137:41755
 TITLE: Antidiabetic agents containing amine derivatives having benzimidazole or imidazopyridine ring and their other uses
 INVENTOR(S): Fujita, Takashi; Wada, Kunio; Oguchi, Minoru; Honma, Eiichi; Fujiwara, Toshihiko
 PATENT ASSIGNEE(S): Sankyo Co., Ltd., Japan
 SOURCE: Jpn. Kokai Tokkyo Koho, 109 pp. CODEN: JKKXAF
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

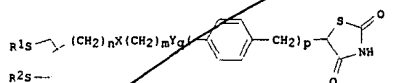
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2002179568	A2	20020626	JP 2001-308814	20011004
PRIORITY APPLN. INFO.:			JP 2000-307159	A 20001006
OTHER SOURCE(S):		MARPAT 137:41755		



AB Prophylactic and/or therapeutic agents for diabetes, glucose intolerance, diabetic complications, or gestational diabetes contain the derivs. I (R1 = carbamoyl which may have 1-2 .alpha., thiocarbamoyl which may have 1-2 .alpha., sulfonyl having 1 .alpha., carbonyl having 1 .alpha.; R2, R3 = H, C1-10 alkyl, C6-10 aryl, which may have 1-3 .beta., C7-16 aralkyl which may have 1-3 .beta. on the aryl moiety; W1-W3 = direct bond, C1-8 alkylene; X, Y, Q = O, S; Z = CH, N; Ar = benzene or naphthalene ring substituted with 1-4 L; L = H, C1-6 alkyl, C6-10 aryl which may have 1-3 .beta., C7-16 aralkyl which may have 1-3 .beta. on the aryl moiety; definitions of .alpha. and .beta. are given) or their pharmacol. acceptable salts. I and their salts are also useful as insulin resistance

L24 ANSWER 2 OF 5 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2002:165043 CAPLUS
 DOCUMENT NUMBER: 136:216739
 TITLE: Preparation of dithiolanyl thiazolidinediones
 AS peroxisome proliferator-activated receptor .gamma. activators.
 INVENTOR(S): Pershadsingh, Harrihar A.; Avery, Mitchell A.
 PATENT ASSIGNEE(S): University of Mississippi, USA
 SOURCE: U.S., 38 pp., Cont.-in-part of U. S. Ser. No. 6,204,288. CODEN: USXXAM
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 6353011	B1	20020305	US 2000-520208	20000307
US 6127394	A	20001003	US 1999-264370	19990308
US 6204288	B1	20010320	US 2000-497324	20000203
PRIORITY APPLN. INFO.:			US 1999-264370	A2 19990308
OTHER SOURCE(S):		MARPAT 136:216739	US 2000-497324	A2 20000203



AB Title compds. [I; R1 R2 = H, COR6 CSR6; R6 = H, alkyl, aryl, aralkyl, carboxy, NR7R8, OR7, NHR7, SR7, NR7R8; R7, R8 = H, alkyl, aryl arylalkyl; R1R2 = atoms to form a 1,2-dithiolane ring; X = O, NR, CO2, OCO2, CONR; R = H, (substituted) alkyl, aryl; Y = O, S, NR3; R3 = H, (substituted) alkyl; n = 2-14; m = 0-14; q, t = 0, 1; when m = 0 then q = 0], were prepd. as peroxisome proliferator-activated receptor .gamma. activators [no data]. Thus, 5-[[4-(2-aminoethoxy)phenyl]methyl]-1,3-thiazolidine-2,4-dione hydrochloride in CH2Cl2 at 0.degree. was treated with Et3N and then with a mixt. prepd. from DL-lipoic acid, Et3N, and iso-Pr chloroformate in

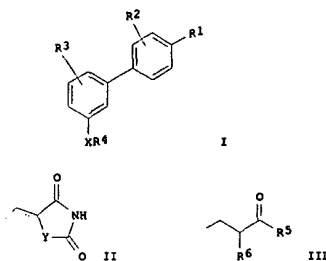
L24 ANSWER 1 OF 5 CAPLUS COPYRIGHT 2002 ACS (Continued)
 improving agents, hypoglycemics, inflammation inhibitors, immunomodulators, aldose reductase inhibitors, 5-lipoxygenase inhibitors, lipid peroxide formation inhibitors, PPAR activators, antiosteoporotic agents, leukotriene antagonists, adipocyte conversion promoters, cancer cell growth inhibitors, and Ca blockers. Feeding diabetic KK mice with feed contg. 0.01% 1-(4-chlorophenyl)-3-[4-[2-[4-(2,4-dioxothiazolidin-5-ylmethyl)phenoxy]methyl]-1-methyl-1H-benzimidazol-6-yloxy]-2,6-dimethylphenyl]thiourea (II) for 3 days showed 48.9% hypoglycemic effect. Capsules, tablets, and granules contg. II were also formulated.

L24 ANSWER 2 OF 5 CAPLUS COPYRIGHT 2002 ACS (Continued)
 toluene/CH2Cl2 followed by stirring to give 56% N-[2-[4-[(2,4-dioxo-1,3-thiazolidin-5-yl)methyl]phenoxy]ethyl]-5-(1,2-dithiolan-3-yl)pentanamide .
 REFERENCE COUNT: 28 THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 3 OF 5 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2002:122976 CAPLUS
 DOCUMENT NUMBER: 136:167181
 TITLE: Preparation of biphenyl derivatives and their use as
 INVENTOR(S): PPAR gamma receptor agonists
 Bernardon, Jean-Michel; Clary, Laurence
 PATENT ASSIGNEE(S): Galderma Research & Development, Fr.
 SOURCE: PCT Int. Appl., 113 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: French
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002012210	A1	20020214	WO 2001-FR2543	20010803
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MY, NZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, SI, KS, KZ, MD, RU, TJ, TM, RW: GR, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
FR 2812876	A1	20020215	FR 2000-10447	20000808
AU 2001085981	A5	20020218	AU 2001-85981	20010803
PRIORITY APPL. INFO.:		FR 2000-10447 A 20000808 WO 2001-FR2543 W 20010803		
OTHER SOURCE(S):		MARPAT 136:167181		
GI				

L24 ANSWER 3 OF 5 CAPLUS COPYRIGHT 2002 ACS (Continued)

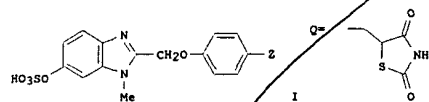


AB The invention concerns compds. I (e.g. N-[[4'-(2,4-dioxothiazolidin-5-ylmethyl)biphenyl-3-yl]methyl]-N-methylbenzamide) wherein: R1 represents a radical II or III; Y represents a CH2 radical or a S atom; R5 represents hydroxy, alkoxy, NH-OH, or N(R8)(R9) radical, and R6 represents alkyl, OR10, SR10, or (CH2)r-COR11. Said compds. are useful as PPAR gamma receptor activators in pharmaceutical compns. for use in human or veterinary medicine (in dermatol., as well as in the field of cardiovascular diseases, immune diseases and/or diseases related to lipid metab.), or in cosmetic compns. Agonist activity for 15 of the claimed compds. is reported. Although the methods of prepn. are not claimed, 82 example preps. are included. REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L24 ANSWER 4 OF 5 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2001:254872 CAPLUS
 DOCUMENT NUMBER: 134:280842
 TITLE: Preparation of 2-phenoxyethyl-1-methyl-6-sulfoxybenzimidazoles with blood sugar-lowering activity
 INVENTOR(S): Iwabuchi, Haruo; Fujiwara, Toshihiko; Fujita, Takashi
 PATENT ASSIGNEE(S): Sankyo Co., Ltd., Japan
 SOURCE: Jpn. Kokai Tokkyo Koho, 28 pp.
 CODEN: JKXKAF
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2001097954	A2	20010410	JP 2000-228085	20000728
PRIORITY APPL. INFO.:		JP 1999-215140 A 19990729		
OTHER SOURCE(S):		MARPAT 134:280842		
GI				

L24 ANSWER 4 OF 5 CAPLUS COPYRIGHT 2002 ACS (Continued)
 (prepn. given) in 100 mL pyridine over a period of 30 min and left to stand at room temp. for 18 h to give. after workup and reversed phase chromatog., 5-[4-(6-sulfoxy-1-methyl-1H-benzimidazol-2-ylmethoxy)benzyl]thiazolidine-2,4-dione (II). A dispersant, tablet, or capsule formulation contg. II was prepd.

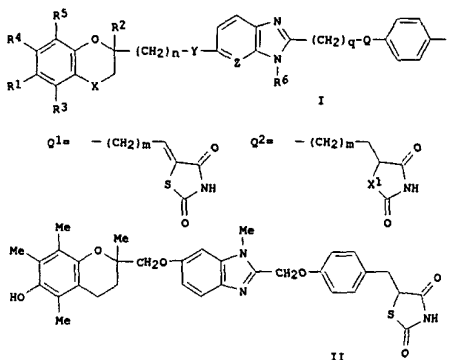


AB The title compds. [I: Z=Q, CH2CH(COY)S(O)nR; wherein Y = OH, NH2; R = C1-5 linear or branched alkyl; n = 0,1,2] are prepd. These compds. also possess insulin-resistance improving, antiinflammatory, immunomodulating, aldose reductase-inhibitory, 5-lipoxygenase-inhibitory, lipid peroxide formation-inhibitory, peroxisome proliferator-activated receptor (PPAR)-activating, anti-osteoporosis, leukotriene antagonist, fat cellularization, cancer proliferation-inhibitory, or calcium antagonist activity and are useful for the prevention or treatment of diseases caused by insulin resistance, diabetes, hyperglycemia, diabetes complications, or cancer (no data). Thus, a soln. of 0.80 mL chlorosulfonic acid in 100 mL MeCN was added dropwise to a soln. of 3.36 g 5-[4-(6-hydroxy-1-methyl-1H-benzimidazol-2-ylmethoxy)benzyl]thiazolidine-2,4-dione hydrochloride

L24 ANSWER 5 OF 5 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2000:742095 CAPLUS
 DOCUMENT NUMBER: 133:296438
 TITLE: Preparation of substituted fused imidazole derivatives
 as hypoglycemics
 INVENTOR(S): Fujita, Takashi; Wada, Kunio; Oguchi, Minoru; Honma, Hidehito; Fujiwara, Toshihiko
 PATENT ASSIGNEE(S): Sankyo Company, Limited, Japan
 SOURCE: PCT Int. Appl., 274 pp.
 CODEN: PIMXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000061582	A1	20001019	WO 2000-JP2217	20000406
W: AU, BR, CA, CN, CZ, HU, ID, IL, IN, KR, MX, NO, NZ, PL, RU, TR, US, ZA				
RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
JP 2000351777	A2	20001219	JP 2000-105985	20000407
PRIORITY APPLN. INFO.: JP 1999-101369 A 19990408				
OTHER SOURCE(S): MARPAT 133:296438				
GI				

L24 ANSWER 5 OF 5 CAPLUS COPYRIGHT 2002 ACS (Continued)



AB Comps. represented by general formula (I) and salts and esters thereof
 [wherein R1 is hydrogen, C1-6 alkyl, (un)substituted C6-10 aryl or C7-16 aralkyl, HO, (un)substituted acyloxy, C1-6 alkoxy, (un)substituted NH2, etc.; R2 is hydrogen, C1-6 alkyl, or (un)substituted C7-16 aralkyl; R4, R4, or R5 is each hydrogen, C1-6 alkyl, or C1-6 alkoxy; R6 is hydrogen, C1-6 alkyl, (un)substituted C6-10 aryl or C7-16 aralkyl; Q and Y are each oxygen or sulfur; X is CH2, CO, CH(OR9), or C(:NOR10); wherein R9 or R10 is hydrogen, (un)substituted C1-6 alkyl, C7-16 aralkyl, or acyl; Z is CH or nitrogen; n and q are each 1 to 5; and A is a group represented by general formula Q1, Q2, Q3, or (CH2)m CH(CO2H)-BR7; wherein m is 0 to 8;
 X1 is oxygen or sulfur; B is oxygen, sulfur, or (un)substituted NH; and R7 is hydrogen, C1-6 alkyl, (un)substituted C6-10 aryl or C7-16 aralkyl, or haloalkyl] are prepd. These comps. are useful as insulin resistance

L24 ANSWER 5 OF 5 CAPLUS COPYRIGHT 2002 ACS (Continued)
 improvers, hypoglycemics, antiinflammatory agents, immunomodulators,
 aldose reductase inhibitors, 5-lipoxygenase inhibitors, lipid peroxide-formation inhibitors, peroxisome proliferator-activated receptor (PPAR) activators, anti-osteoporosis agents, leukotriene antagonists, promoters of fat cell formation, cancer cell-proliferation inhibitors, or calcium antagonists. They are useful for the prevention or treatment of diabetes, hyperlipidemia, obesity, glucose tolerance insufficiency, hypertension, fatty liver, diabetes complication, arteriosclerosis, gestational diabetes, polycystic ovarian syndrome, cardiovascular diseases, cell damages caused by atherosclerosis or ischemic heart diseases, gout, osteoarthritis, rheumatic arthritis, allergic diseases, asthma, gastrointestinal ulcer, cachexia, autoimmune diseases, cancer, osteoporosis, or cataract. Thus, N-[2-amino-5-(6-methoxymethoxy-2,5,7,8-tetramethylchroman-2-ylmethoxy)phenyl]-N-methylcarbamoyl acid tert-Bu ester was condensed with 4-(2,4-dioxothiazolidin-5-ylmethyl)phenoxyacetic acid using di-Et cyanophosphate and Et3N in THF at room temp. for 30 min, followed by treatment of the product with 4 N HCl/dioxane at room temp. for 5 h gave 5-[4-(6-(6-hydroxy-2,5,7,8-tetramethylchroman-2-ylmethoxy)-1-methyl-1H-benzimidazol-2-ylmethoxy)benzyl]thiazolidine-2,4-dione hydrochloride (II.HCl). When a feed contg. 0.01% II.HCl was fed to mice for 3 days, the blood sugar level was lowered by 66.7% compared to control animal.
 REFERENCE COUNT: 18 THERE ARE 18 CITED REFERENCES AVAILABLE FOR THIS
 RECORD. ALL CITATIONS AVAILABLE IN THE
 RE FORMAT

=> log y

COST IN U.S. DOLLARS

SINCE FILE	TOTAL
ENTRY	SESSION
24.12	438.69

FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE	TOTAL
ENTRY	SESSION
-4.34	-40.89

CA SUBSCRIBER PRICE

STN INTERNATIONAL LOGOFF AT 10:17:52 ON 12 AUG 2002

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID:sssptal610jxm

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

08.12.02
STN + Author
Search.

* * * * * Welcome to STN International * * * * *

NEWS 1 Web Page URLs for STN Seminar Schedule - N. America
NEWS 2 Apr 08 "Ask CAS" for self-help around the clock
NEWS 3 Apr 09 BEILSTEIN: Reload and Implementation of a New Subject Area
NEWS 4 Apr 09 ZDB will be removed from STN
NEWS 5 Apr 19 US Patent Applications available in IFICDB, IFIPAT, and
IFIUDB
NEWS 6 Apr 22 Records from IP.com available in CAPLUS, HCAPLUS, and
ZCAPLUS
NEWS 7 Apr 22 BIOSIS Gene Names now available in TOXCENTER
NEWS 8 Apr 22 Federal Research in Progress (FEDRIP) now available
NEWS 9 Jun 03 New e-mail delivery for search results now available
NEWS 10 Jun 10 MEDLINE Reload
NEWS 11 Jun 10 PCTFULL has been reloaded
NEWS 12 Jul 02 FOREGE no longer contains STANDARDS file segment
NEWS 13 Jul 22 USAN to be reloaded July 28, 2002;
saved answer sets no longer valid
NEWS 14 Jul 29 Enhanced polymer searching in REGISTRY
NEWS 15 Jul 30 NETFIRST to be removed from STN
NEWS 16 Aug 08 CANCERLIT reload
NEWS 17 Aug 08 PHARMAMarketLetter(PHARMAML) - new on STN
NEWS 18 Aug 08 NTIS has been reloaded and enhanced
NEWS 19 Aug 09 JAPIO to be reloaded August 18, 2002

NEWS EXPRESS February 1 CURRENT WINDOWS VERSION IS V6.0d,
CURRENT MACINTOSH VERSION IS V6.0a(ENG) AND V6.0Ja(JP),
AND CURRENT DISCOVER FILE IS DATED 05 FEBRUARY 2002
NEWS HOURS STN Operating Hours Plus Help Desk Availability
NEWS INTER General Internet Information
NEWS LOGIN Welcome Banner and News Items
NEWS PHONE Direct Dial and Telecommunication Network Access to STN
NEWS WWW CAS World Wide Web Site (general information)

Enter NEWS followed by the item number or name to see news on that
specific topic.

All use of STN is subject to the provisions of the STN Customer
agreement. Please note that this agreement limits use to scientific
research. Use for software development or design or implementation
of commercial gateways or other similar uses is prohibited and may
result in loss of user privileges and other penalties.

* * * * * STN Columbus * * * * *

FILE 'HOME' ENTERED AT 09:59:58 ON 12 AUG 2002

=> fil reg

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

0.42

0.42

FILE 'REGISTRY' ENTERED AT 10:01:05 ON 12 AUG 2002

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2002 American Chemical Society (ACS)

STRUCTURE FILE UPDATES: 9 AUG 2002 HIGHEST RN 443534-23-4

DICTIONARY FILE UPDATES: 9 AUG 2002 HIGHEST RN 443534-23-4

TSCA INFORMATION NOW CURRENT THROUGH January 7, 2002

Please note that search-term pricing does apply when
conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Calculated physical property data is now available. See HELP PROPERTIES
for more information. See STN Note 27, Searching Properties in the CAS
Registry File, for complete details:

<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

=>

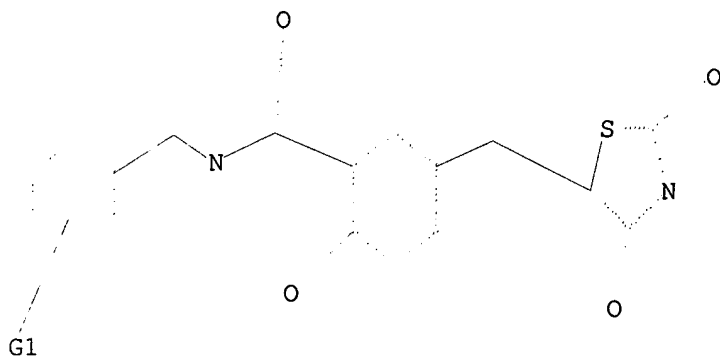
Uploading 10049937 prlim.str

L1 STRUCTURE UPLOADED

=> d

L1 HAS NO ANSWERS

L1 STR



C1 2
H 1

G1 [01], [02]

Structure attributes must be viewed using STN Express query preparation.

=> s l1 ful
FULL SEARCH INITIATED 10:02:19 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 108 TO ITERATE

100.0% PROCESSED 108 ITERATIONS
SEARCH TIME: 00.00.01

46 ANSWERS

L2 46 SEA SSS FUL L1

=> s l2 and caplus/lc
23846251 CAPLUS/LC
L3 46 L2 AND CAPLUS/LC

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	146.18	146.60

FILE 'CAPLUS' ENTERED AT 10:04:24 ON 12 AUG 2002
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2002 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 12 Aug 2002 VOL 137 ISS 7
FILE LAST UPDATED: 11 Aug 2002 (20020811/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

CAS roles have been modified effective December 16, 2001. Please check your SDI profiles to see if they need to be revised. For information on CAS roles, enter HELP ROLES at an arrow prompt or use the CAS Roles thesaurus (/RL field) in this file.

=> s l2 ful
L4 32 L2

=> s blood?
L5 990331 BLOOD?

=> s l4 and l5
L6 12 L4 AND L5

=> d l4 1-32 ibib abs hitstr

L4 ANSWER 1 OF 32 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2002:540258 CAPLUS
TITLE: Preparation of benzoxepinopyridines as HMG-CoA
reductase inhibitors
INVENTOR(S): Robl, Jeffrey A.; Chen, Bang-chi; Sun,
Chong-qing
PATENT ASSIGNEE(S): USA
SOURCE: U.S. Pat. Appl. Publ., 42 pp., Cont.-in-part
of U.S.
DOCUMENT TYPE: Ser. No. 875,155.
LANGUAGE: CODEN: USXXCO
FAMILY ACC. NUM. COUNT: 2 Patent
English
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2002094977	A1	20020718	US 2001-7407	20011204
US 2002013334	A1	20020131	US 2001-875155	20010606

PRIORITY APPLN. INFO.: US 2000-211595P P 20000615
US 2001-875155 A2 20010606

GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE
PRINT *

AB Title compds. I [X = O, S, SO, SO₂, NR₇; Z = HOCHCH₂CH(OH)CH₂CO₂R₃,
4-hydroxy-2-oxopyran-6-yl, etc.; n = 0, 1; R₁, R₂ = alkyl,
arylalkyl,
cycloalkyl, alkenyl, cycloalkenyl, aryl, heteroaryl,
cycloheteroalkyl; R₃
= H, alkyl, metal ion; R₄ = H, halo, CF₃, etc.; R₇ = H, alkyl,
aryl,
alkanoyl, aroyl, alkoxycarbonyl, etc.; R₉, R₁₀ = H, alkyl], were
prepd. as
HMG CoA reductase inhibitors active in inhibiting cholesterol
biosynthesis, modulating blood serum lipids such as lowering LDL
cholesterol and/or increasing HDL cholesterol, and treating
hyperlipidemia, hypercholesterolemia, hypertriglyceridemia and
atherosclerosis (no data). E.g., a multistep synthesis of II is
reported.

IT 213252-19-8, KRP297
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
(Biological study); USES (Uses)
(coadministered agents; prepn. of benzoxepinopyridines as

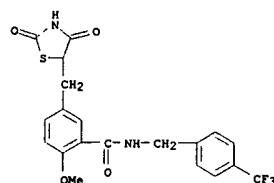
HMG-CoA
reductase inhibitors for the treatment of hyperlipidemia,
hypercholesterolemia, hypertriglyceridemia, atherosclerosis,
and other

disorders)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-
(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 1 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



L4 ANSWER 2 OF 32 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2002:504648 CAPLUS
DOCUMENT NUMBER: 137:83637
TITLE: Medicinal compositions containing diuretic and
insulin
INVENTOR(S): resistance-improving agent
Takaka, Masaya; Araki, Kazushi; Kanda, Shoichi
PATENT ASSIGNEE(S): Sankyo Company, Limited, Japan
SOURCE: PCT Int. Appl., 183 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002051441	A1	20020704	WO 2001-JP11296	20011221

PH, PL, W: AU, BR, CA, CN, CO, CZ, HU, ID, IL, IN, KR, MX, NO, NZ,
RU, SG, SK, US, VN, ZA
RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU,
MC, NL, PT, SE, TR

PRIORITY APPLN. INFO.: JP 2000-394424 A 20001226

OTHER SOURCE(S): MARPAT 137:83637

AB Disclosed are medicinal compns. contg. a diuretic and an insulin
resistance-improving agent whereby side effects assocg. the
administration
of an insulin resistance-improving agent (for example,
megalocardia,
edema, body fluid retention, pleural effusion) can be prevented or
treated. Oral administration of furosemide prevented increases of
heart

wt. and blood plasma, and edema due to administration of

5-[(4-(6-methoxy-1-methyl-1H-benzimidazol-2-ylmethoxy)benzyl]thiazolidine

2,4-dione hydrochloride.

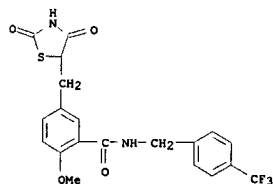
IT 213252-19-8, KRP-297

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(medicinal compns. contg. diuretics and insulin
resistance-improving
agents)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-
(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 2 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



REFERENCE COUNT: 22 THERE ARE 22 CITED REFERENCES AVAILABLE
FOR THIS
RE FORMAT RECORD. ALL CITATIONS AVAILABLE IN THE

L4 ANSWER 3 OF 32 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2002:409256 CAPLUS
 DOCUMENT NUMBER: 137:735
 TITLE: Methods and compositions for treatment of diabetes and related conditions via gene therapy
 INVENTOR(S): Caplan, Shari L.; Boettcher, Brian R.; Slosberg, Eric
 D.: Connelly, Sheila; Kaleko, Michael; Desai, Urvi J.
 PATENT ASSIGNEE(S): USA
 SOURCE: U.S. Pat. Appl. Publ., 42 pp.
 CODEN: USXXCO
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

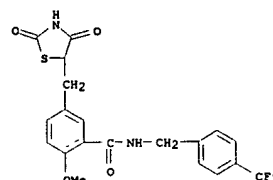
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2002065239	A1	20020530	US 2001-808457	20010314
PRIORITY APPLN. INFO.: US 2000-266328P P 20000315				

AB Methods and compns. are disclosed for the treatment of diabetes, obesity and diabetic-related conditions. The methods include gene therapy based administration of a therapeutically effective amt. of vectors encoding the following: glucokinase regulatory protein alone or co-administered with glucokinase or with metab. modifying proteins; glucokinase co-administered with metab. modifying proteins; or glucokinase regulatory protein co-administered with glucokinase in combination with metab. modifying proteins, to a diabetic patient. The metab. modifying proteins include UCP2, UCP3, PPAR.alpha., OB-Rb, GLP-1 and GLP-1 analogs (administered via vector or directly as a peptide). Preferred examples of GLP-1 analogs include GLP-1-Gly8, Exendin-4 and the "Black Widow" chimeric GLP-1 analog. Addnl., PPAR.alpha. ligands and DPP-IV inhibitors may be co-administered with the above.

IT 213252-19-8, KRP-297
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (gene therapy for treatment of diabetes and related conditions)

RN 213252-19-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)

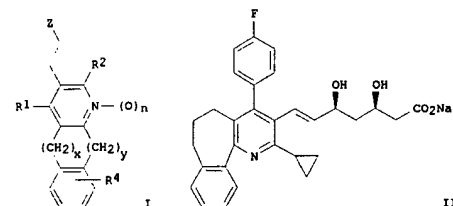
L4 ANSWER 3 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



L4 ANSWER 4 OF 32 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2002:392237 CAPLUS
 DOCUMENT NUMBER: 136:401651
 TITLE: Preparation of fused pyridine derivatives as reductase inhibitors
 INVENTOR(S): Robl, Jeffrey A.; Chen, Bang-Chi; Sun, Chong-Qing
 PATENT ASSIGNEE(S): USA
 SOURCE: U.S. Pat. Appl. Publ., 46 pp., Cont.-in-part of U.S.
 Ser. No. 875,218.
 CODEN: USXXCO
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2002061901	A1	20020523	US 2001-8154	20011204
US 2002028826	A1	20020307	US 2001-875218	20010606
PRIORITY APPLN. INFO.: US 2000-211594P P 20000615				
US 2001-875218 A2 20010606				

OTHER SOURCE(S): MARPAT 136:401651
 GI

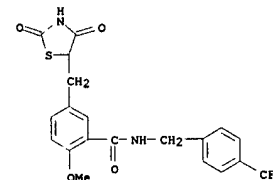


AB The title compds. I and their pharmaceutically acceptable salts, esters, prodrug esters, and stereoisomers are claimed (wherein: Z = CH(OH)CH2CR7(OH)CH2CO2R3 or corresponding pyranone lactone derivs.; n = 0, 1; x = 0, 1, 2, 3, or 4; y = 0, 1, 2, 3 or 4, provided that at least one of x and y is other than 0; and optionally one or more carbons of (CH2)x and/or (CH2)y together with addnl. carbons form a 3 to 7 membered spirocyclic ring; R1, R2 = alkyl, arylalkyl, cycloalkyl, alkenyl, cycloalkenyl, aryl, heteroaryl, cycloheteroalkyl; R3 = H or lower alkyl;

L4 ANSWER 4 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
 R4 = H, halo, CF3, OH, alkyl, alkoxy, CO2H, (un)substituted NH2, cyano, (un)substituted CONH2, etc.; R7 = H, alkyl. The compds. are HMG-CoA reductase inhibitors, and are active in inhibiting cholesterol biosynthesis and modulating blood serum lipids, for example, lowering LDL cholesterol and/or increasing HDL cholesterol (no data). I are thus useful in treating hyperlipidemia and dyslipidemia, in hormone replacement therapy, and in treating hypercholesterolemia, hypertriglyceridemia and atherosclerosis, as well as Alzheimer's disease and osteoporosis. Prepn. of several compds. are described. For instance, a multistep synthesis of fused pyridine deriv. II is reported. Compds. I may be used in a manner similar to atorvastatin, pravastatin, simvastatin, etc. Combinations of compds. I with various other drugs are claimed, the latter being specified as certain pharmacol. classes, as inhibitors of specific enzymes, as (ant)agonists of specific receptors, and as numerous named drugs.

IT 213252-19-8, KRP297
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (therapeutic compns. also contg.; prepn. of fused pyridine derivs. as HMG-CoA reductase inhibitors)

RN 213252-19-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 5 OF 32 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2002:142553 CAPLUS
DOCUMENT NUMBER: 136:177969
TITLE: Medicinal compositions containing PPAR.gamma.
agonists
and RXR agonists for preventing and treating
cancer
INVENTOR(S): Kurakata, Shinichi; Fujiwara, Kosaku;
Shimazaki,
Naomi; Fujita, Takashi
PATENT ASSIGNEE(S): Sankyo Company, Limited, Japan
SOURCE: PCT Int. Appl., 40 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002013864	A1	20020221	WO 2001-JP7037	20010815
W: AU, BR, CA, CN, CO, CZ, HU, ID, IL, IN, KR, MX, NO, NZ, PL, RU, SG, SK, US, ZA				
RM: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR				
JP 2002128700	A2	20020509	JP 2001-241740	20010809
AU 2001078738	A5	20020225	AU 2001-78738	20010815
PRIORITY APPLN. INFO.: JP 2000-246910 A 20000816 JP 2000-2000246910A 20000816 WO 2001-JP7037 W 20010815				

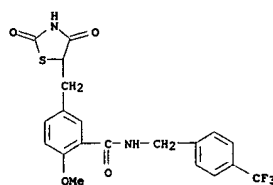
OTHER SOURCE(S): MARPAT 136:177969
AB Disclosed are medicinal compns. for preventing or treating cancer
wherein
one or more Peroxisome proliferator-activated receptor .gamma.
(PPAR.gamma.) activation agonists and one or more retinoid X
receptor
(RXR) activation agonists are used simultaneously or successively.
A
combined administration of 5-[4-(6-methoxy-1-methylbenzimidazol-2-
ylmethoxy)benzyl]thiazolidine-2,4-dione hydrochloride (I) 5 and
targretin
100 mg/kg to HL-60 cell-bearing mice showed synergistic antitumor
effect.
Also, tablets were prepd. from I 0.004, targretin 0.1, lactose
0.244, corn
starch 50, and magnesium stearate 0.002 g.
IT 213252-19-8
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(simultaneous or successive use of PPAR.gamma. agonists and RXR
agonists for prevention or treatment of cancer)
RN 213252-19-8 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-
(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 6 OF 32 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2002:142506 CAPLUS
DOCUMENT NUMBER: 136:177977
TITLE: Methods for treating inflammatory diseases
using PPAR
agonists
INVENTOR(S): Pershadsingh, Narrihar A.
PATENT ASSIGNEE(S): USA
SOURCE: PCT Int. Appl., 42 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 2
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002013812	A1	20020221	WO 2001-US25668	20010816
W: AU, CA, MX, NZ, US				
RM: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR				
AU 2001088271	A5	20020225	AU 2001-88271	20010816
PRIORITY APPLN. INFO.: US 2000-225907P P 20000817 US 2000-230509P P 20000906 WO 2001-US25668 W 20010816				

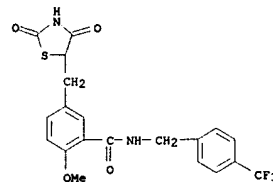
AB The present invention describes methods for the use of PPAR
ligands in the
treatment inflammatory endocrine, dermatol., cardiovascular
immunol.,
neurolog., ophthalmic, neoplastic, pulmonary diseases, and
age-related
dysregulations. In addn., methods are provided for treating said
conditions and diseases comprising the step of administering to a
human or
an animal in need thereof a therapeutic amt. of pharmacol. compns.
comprising a pharmaceutically acceptable carrier, and a PPAR.gamma.
agonist which cross-activates PPAR.alpha. or PPAR.delta. or both,
or a
PPAR.gamma. partial agonist, or a PPAR.gamma./RXR agonist,
effective to
reverse, slow, stop, or prevent the pathol. inflammatory or
degenerative
process.
IT 213252-19-8, KRP 297
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
(Biological study); USES (Uses)
(methods for treating inflammatory diseases using PPAR agonists)
RN 213252-19-8 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-
(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 5 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE
FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE
RE FORMAT

L4 ANSWER 6 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)

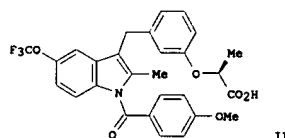
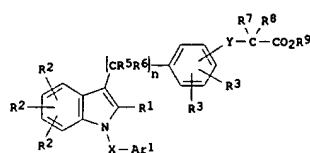


REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE
FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE
RE FORMAT

L4 ANSWER 7 OF 32 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2002:90008 CAPLUS
 DOCUMENT NUMBER: 136:151071
 TITLE: Preparation of N-substituted indoles for treating diabetes
 INVENTOR(S): Acton, John J., III; Black, Regina Marie; Jones,
 Anthony Brian; Wood, Harold Blair
 PATENT ASSIGNEE(S): Merck & Co., Inc., USA
 SOURCE: PCT Int. Appl., 73 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002008188	A1	2002/01/31	WO 2001-US22979	2001/07/20
W: AE, AG, AL, AM, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
US 2002042441	A1	2002/04/11	US 2001-912961	2001/07/25
PRIORITY APPLN. INFO.: US 2000-220778P			P 2000/07/25	
OTHER SOURCE(S): MARPAT 136:151071				

L4 ANSWER 7 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



AB The title indoles having aryloxyacetic acid substituents [I; R1 = Me, optionally substituted with 1-3 F atoms; R2-R4 = H, halo, alkyl, etc.; R5, R6 = H, F, OH, alkyl; and R5 and R6 groups that are on the same carbon atom optionally may be joined to form a cyclopropyl group; R7, R8 = H, F, alkyl; or CR7R8 may form cycloalkyl; R9 = H, alkyl; Ar1 = (un)substituted Ph, naphthyl, pyridyl, quinolyl; X = CO, SO2, CH2, CHMe, CMe2, CF2, cyclopropylidene; Y = O, S; n = 0-5] which are agonists or partial agonists of PPAR gamma, and are useful in the treatment, control or prevention of non-insulin dependent diabetes mellitus (NIDDM), hyperglycemia, dyslipidemia, hyperlipidemia, hypercholesterolemia, hypertriglyceridemia, atherosclerosis, obesity, vascular restenosis, inflammation, and other PPAR mediated diseases, disorders and conditions.

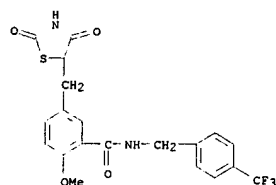
IT 213252-19-8, KRP-297

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (prepn. of N-substituted indoles for treating diabetes)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 7 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



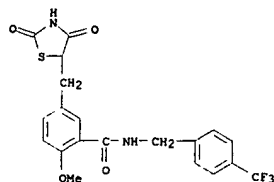
REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 8 OF 32 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2002:56491 CAPLUS
 DOCUMENT NUMBER: 137:73203
 TITLE: Pharmacological analysis of wild-type .alpha., .gamma. and .delta. subtypes of the human peroxisome proliferator-activated receptor
 AUTHOR(S): Wurch, T.; Junquero, D.; Delhon, A.; Pauwels, P. J.
 CORPORATE SOURCE: Department of Cellular and Molecular Biology, de Recherche Pierre Fabre, Castres, 81106, Fr.
 SOURCE: Naunyn-Schmiedeberg's Archives of Pharmacology (2002), 365(2), 133-140
 CODEN: NSAPCC; ISSN: 0028-1298
 PUBLISHER: Springer-Verlag
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Three distinct peroxisome proliferator-activated receptor (PPAR) subtypes were isolated from human brain RNA. Whereas the PPAR.delta. subtype perfectly matched the amino acid sequences reported in the Genbank database, several differences were found for the PPAR.alpha. (Lys123Met, Ala268Val, Gly296Ala and Val444Ala) and PPAR.gamma.2 (Met81Ile, Pro9Ala, Met186Ile, Pro187Ala and the deletion of a Gln213 residue) subtypes. A pharmacol. anal. was undertaken by co-expressing each PPAR subtype with a reporter plasmid contg. a luciferase gene under the transcriptional control of a synthetic, triplicated PPAR response element in either HepG2 or Cos-7 cells. Whereas fenofibrate unselectively activated the PPAR.alpha. and PPAR.delta. subtypes, the related BM-17.0744 compd. was more potent and selective for PPAR.alpha.. The thiazolidine dione derivs. rosiglitazone and pioglitazone were potent and selective PPAR.gamma.2 agonists. L-165041, reported as a selective and potent PPAR.delta. ligand, displayed in this specified transactivation system, apart from its highly efficacious PPAR.delta. agonist activity, partial and full agonism at, resp., PPAR.alpha. and PPAR.gamma.2 subtypes. In conclusion, transcriptional control of a luciferase gene by wild-type PPAR subtypes provides powerful recombinant assays to evaluate ligand's efficacy at these nuclear receptors.

IT 213252-19-8, KRP-297

RL: PAC (Pharmacological activity); BIOL (Biological study) (pharmacol. anal. of wild-type .alpha., .gamma. and .delta. subtypes of

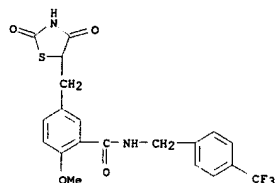
L4 ANSWER 8 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
human peroxisome proliferator-activated receptor)
RN 213252-19-8 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 40 THERE ARE 40 CITED REFERENCES AVAILABLE
FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE
RE FORMAT

L4 ANSWER 9 OF 32 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:90080 CAPLUS
DOCUMENT NUMBER: 136:318816
TITLE: Design, synthesis and evaluation of substituted phenylpropanoic acid derivatives as peroxisome proliferator-activated receptor (PPAR)
activators:
AUTHOR(S): Miyachi, Hiroyuki; Nomura, Masahiro; Tanase, Takahiro;
Masaki;
CORPORATE SOURCE: Murakami, Koji; Awano, Katsuya
Kyorin Pharmaceutical Co., Ltd., Discovery Research Laboratories, Tochigi, Shimotsuga-gun, Nogi-machi,
329-0114, Japan
SOURCE: Bioorganic & Medicinal Chemistry Letters (2001), Volume Date 2002, 12(1), 77-80
CODEN: BMCLE8; ISSN: 0960-894X
PUBLISHER: Elsevier Science Ltd.
DOCUMENT TYPE: Journal
LANGUAGE: English
AB A series of substituted phenylpropanoic acid derivs. was prepd. as part of a search for subtype-selective human peroxisome proliferator-activated receptor (PPAR) activators. Structure-activity relationship studies indicated that the substituent at the .alpha.-position of the carboxyl group plays a key role in detg. the potency and the selectivity for PPAR transactivation.
IT 213252-19-8, KRP 297
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(design, synthesis and evaluation of substituted phenylpropanoic acid derivs. as PPAR activators)
RN 213252-19-8 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 9 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



REFERENCE COUNT: 19 THERE ARE 19 CITED REFERENCES AVAILABLE
FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE
RE FORMAT

L4 ANSWER 10 OF 32 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:798208 CAPLUS
DOCUMENT NUMBER: 135:344474
TITLE: Preparation of novel stable crystal of thiazolidinedione derivative
INVENTOR(S): Oonoda, Michiro; Orita, Kazuo
PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
SOURCE: PCT Int. Appl., 17 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001081327	A1	20011101	WO 2001-JP3450	20010423

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
PRIORITY APPLN. INFO.: JP 2000-124006 A 20000425
AB Claimed is a crystal of 5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]benzamide (KRP-297) having diffraction angles (2.theta.) at at least 9.7.degree., 15.0.degree., and 22.5.degree. in X-ray powder diffractometry. The novel crystal of KRP-297 (a known antidiabetic agent) is prepd. through recrystn. from an alc. solvent.
IT 353275-24-8P
RL: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(prepn. of novel stable crystal of thiazolidinedione deriv.)
RN 353275-24-8 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]-, monosodium salt (9CI) (CA INDEX NAME)

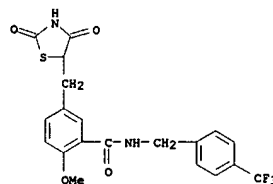
L4 ANSWER 12 OF 32 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:617810 CAPLUS
DOCUMENT NUMBER: 135:175429
TITLE: Modulation of bone formation with peroxisome proliferator-activated receptor activators and ligands
INVENTOR(S): Scutt, Andrew; Still, Karen
PATENT ASSIGNEE(S): University of Sheffield, UK
SOURCE: PCT Int. Appl., 27 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001060355	A1	20010823	WO 2001-GB626	20010215
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
PRIORITY APPLN. INFO.: GB 2000-3310 A 20000215				
AB The use of an activator or ligand of a peroxisome proliferator-activated receptor, other than PPAR.gamma., or pharmaceutically acceptable deriv. of said activator or ligand, in the manuf. of a medicament for the treatment or prophylaxis of bone disease allows, for the first time, bone anabolism to enhance the deposition of bone in conditions which would benefit from increased bone deposition. The reverse, where there is inhibition and/or retardation of bone deposition is also facilitated.				
IT 213252-19-8, KRP-297				
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (modulation of bone formation with peroxisome proliferator-activated receptor activators and ligands)				

L4 ANSWER 13 OF 32 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:581862 CAPLUS
DOCUMENT NUMBER: 135:152800
TITLE: Alkali metal salt of thiazolidine-2,4-dione derivative
INVENTOR(S): and purification of KRP-297
Noriyuki Ohnoda, Michiro; Orita, Kazuo; Yoshida,
PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
SOURCE: PCT Int. Appl., 17 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

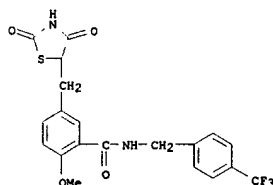
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001057007	A1	20010809	WO 2001-JP598	20010130
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
PRIORITY APPLN. INFO.: JP 2000-23610 A 20000201				
AB This document discloses a method of industrially advantageously purifying 5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxy-N-[(4-(trifluoromethyl)phenyl)methyl]benzamide (KRP-297), a known antidiabetic agent. The method comprises the steps of: forming an alkali metal salt of KRP-297 and a hydrate thereof in a reaction for producing KRP-297; isolating and purifying them; and then liberating the KRP-297 from the salt. Also provided are an alkali metal salt of KRP-297 and a hydrate of the salt.				
IT 213252-19-8P, KRP 297				
RL: PUR (Purification or recovery); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (purifn. of antidiabetic KRP-297)				
RN 213252-19-8 CAPLUS				

L4 ANSWER 12 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
RN 213252-19-8 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)

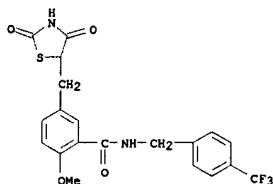


REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

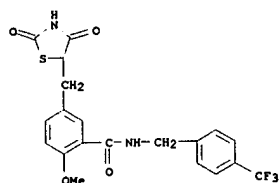
L4 ANSWER 13 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)



IT 353275-24-8P 353275-26-0P 353275-27-1P
353275-28-2P
RL: SPN (Synthetic preparation); PREP (Preparation) (purifn. of antidiabetic KRP-297)
RN 353275-24-8 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-(trifluoromethyl)phenyl)methyl]-, monosodium salt (9CI) (CA INDEX NAME)



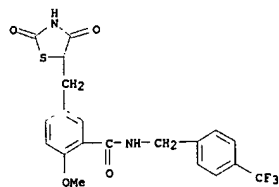
RN 353275-26-0 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-(trifluoromethyl)phenyl)methyl]-, monosodium salt, monohydrate (9CI) (CA INDEX NAME)



● Na

● H₂O

RN 353275-27-1 CAPLUS
 CN Benamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]-, monopotassium salt (9CI) (CA INDEX NAME)



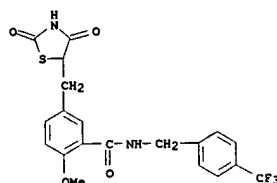
● K

RN 353275-28-2 CAPLUS
 CN Benamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]-, monopotassium salt, monohydrate (9CI) (CA INDEX NAME)

L4 ANSWER 14 OF 32 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2001:564869 CAPLUS
 DOCUMENT NUMBER: 135:132451
 TITLE: Novel remedies with the use of .beta.3 agonists
 INVENTOR(S): Ogawa, Kohei; Umeno, Hiroshi
 PATENT ASSIGNEE(S): Asahi Kasei K. K., Japan
 SOURCE: PCT Int. Appl., 49 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001054728	A1	20010802	WO 2001-JP553	20010126
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				

PRIORITY APPLN. INFO.: JP 2000-20733 A 20000128
 AB Remedies contg. at least one member selected from the group consisting of cholinolytics, monoamine reuptake inhibitors, lipase inhibitors, selective serotonin reuptake inhibitors, insulin, insulin secretion promoters, biguanide, .alpha.-glucosidase inhibitors, insulin resistance improving agents, HMC-CoA reductase inhibitors, anion exchange resins, clofibrate-base drugs and nicotinic acid-base drugs and a compd. having a .beta.3-agonistic activity. The .beta.3 agonist has an activity of inhibiting urination disorder. When used together with a remedy for urination disorder such as propiverine, oxybutynin hydrochloride or tolterodine, it exerts an enhanced anti-urination disorder effect. When used together with an antiobesity agent such as sibutramine or orlistat, it exerts an enhanced antiobesity effect. When used together with an antidiabetic agent such as insulin, glibenclamide, acarbose or rosiglitazone, it exerts an enhanced antidiabetic effect. When used

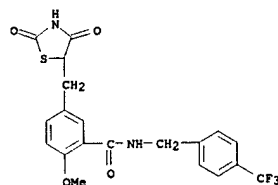


● K

● H₂O

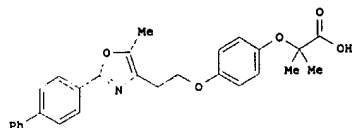
REFERENCE COUNT: 33 THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 14 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
 together with an antilipemic drug such as bezafibrate or pravastatin, it exerts an enhanced antilipemic effect.
 IT 213252-19-8, KRP 297
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (novel remedies with the use of .beta.3 agonists as antidiabetics and antilipidemics and for treatment of urination disorder)
 RN 213252-19-8 CAPLUS
 CN Benamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 43 THERE ARE 43 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 15 OF 32 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2001:367156 CAPLUS
 DOCUMENT NUMBER: 135:131731
 TITLE: Design and Synthesis of
 2-Methyl-2-[4-[2-(5-methyl-2-
 arylloxazol-4-yl)ethoxy]phenoxy]propionic
 Acids: A New Class of Dual PPAR.alpha./gamma. Agonists
 AUTHOR(S): Brooks, Dawn A.; Etgen, Garret J.; Rito,
 Christopher J.; Shuker, Anthony J.; Dominianni, Samuel J.;
 Paterniti, James Warshawsky, Alan M.; Ardecky, Robert;
 R.; Tyhonas, John; Karanewsky, Donald S.;
 Kauffman, Raymond F.; Broderick, Carol L.; Oldham, Brian
 A.;
 Leonard L.: Montrose-Rafizadeh, Chahzrad; Winneroski,
 CORPORATE SOURCE: Faul, Margaret M.; McCarthy, James R.
 Lilly Lilly Research Laboratories A Division of Eli
 Company Lilly Corporate Center, Indianapolis,
 IN, 46285, USA
 SOURCE: Journal of Medicinal Chemistry (2001), 44(13),
 2061-2064
 CODEN: JMCHAR; ISSN: 0022-2623
 PUBLISHER: American Chemical Society
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 GI

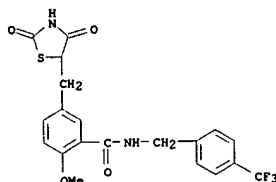


AB Propionic acid deriv. I, which was designed and synthesized based
 on putative pharmacophores of known PPAR.gamma.- and
 PPAR.alpha.-selective compds., exhibits potent dual PPAR.alpha./gamma. agonist activity
 as demonstrated by in vitro binding and dose overlap in the newly
 introduced EOB mouse model for glucose lowering and lipid/cholesterol
 homeostasis.
 IT 213252-19-8, KRP-297

L4 ANSWER 16 OF 32 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2001:359797 CAPLUS
 DOCUMENT NUMBER: 134:344620
 TITLE: Solid oral composition containing KRP-297
 INVENTOR(S): Ohyama, Toshinori; Imamizu, Masaru
 PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 11 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

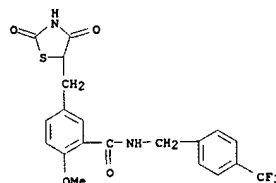
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001034148	A1	20010517	WO 2000-JP7905	20001110
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN,				
CR, CU,	CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,			
ID, IL,	IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,			
LV, MD,	MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG,			
SI, SK,	SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM,			
AZ, BY,	KG, KZ, MD, RU, TJ, TM			
	RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE,			
CH, CY,	DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,			
TR, BF,	BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
PRIORITY APPLM. INFO.: JP 1999-320586 A 19991111				
AB Disclosed are solid compns. for oral use for facilitating the administration in a small dose of KRP-297, which is a ligand common to peroxisome proliferator-activated receptors PPAR.alpha. and PPAR.gamma. (i.e., nuclear receptors) and has an effect of ameliorating insulin resistance, which contain the drug ingredient in a uniform content and can be easily and quant. taken. By prepg. solid compns. for oral use composed of a trace amt. of the drug ingredient together with pharmaceutical carriers, it is possible to provide tablets which contain the drug component in a uniform content and can be easily and quant. taken.				
A film-coated tablet was prepd. from KRP-297 0.25, lactose 78.55, cryst. cellulose 26.2, low-substituted hydroxypropyl cellulose 12, polyvinyl alc. 2.4, magnesium stearate 0.6, hydroxypropyl Me cellulose, and carnauba wax 0.001 mg.				
IT 213252-19-8, KRP-297 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (solid oral compns. contg. uniform contents of KRP-297)				

L4 ANSWER 15 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
 RL: BAC (Biological activity or effector, except adverse); BSU
 (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological
 study); USES (Uses)
 (design and synthesis of
 2-methyl-2-[4-[2-(5-methyl-2-arylloxazol-
 4-yl)ethoxy]phenoxy]propionic acids: a new class of dual
 PPAR.alpha./gamma. agonists)
 RN 213252-19-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-
 (trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 24 THERE ARE 24 CITED REFERENCES AVAILABLE
 FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE
 RE FORMAT

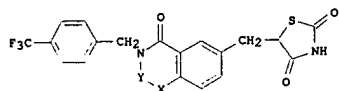
L4 ANSWER 16 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
 RN 213252-19-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-
 (trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE
 FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE
 RE FORMAT

L4 ANSWER 17 OF 32 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:347100 CAPLUS
DOCUMENT NUMBER: 134:353303
TITLE: preparation of thiazolidinyl-containing
bicyclic
heterocycles as humane peroxisome proliferator-
activated receptor .gamma. agonists
INVENTOR(S): Nomura, Masahiro; Murakami, Koji; Kakuta,
Masaki
PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
SOURCE: Jpn. Kokai Tokkyo Koho, 7 pp.
CODEN: JKKXAF
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2001131173	A2	20010515	JP 2000-242708	20000810
PRIORITY APPLN. INFO.: OTHER SOURCE(S):			JP 1999-235531 A	19990823
GI			MARPAT 134:353303	

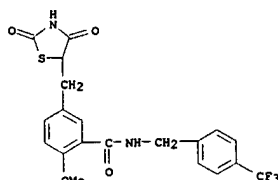


AB Title compds. I (YX = CO₂, CH₂O, CH:CH), their pharmaceutically acceptable salts, or hydrates, useful as for treatment of Type II diabetes and hyperlipemia, are prepd.
2-Hydroxy-5-[(2,4-dioxothiazolidin-5-yl)methyl]-N-[(4-trifluorophenyl)methyl]benzamide was reacted with trioxane in the presence of AcOH in CH₂Cl₂ at room temp. for 2 day to give 421

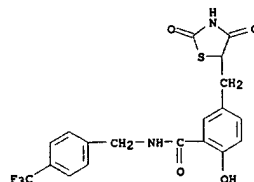
6-[(2,4-dioxothiazolidin-5-yl)methyl]-3-[(4-trifluorophenyl)methyl]-1,3-benzoxazin-4-one showing good transcription activity of proliferator-activated receptor .gamma. in vitro.

IT 213252-19-8
RL: RCT (Reactant); RACT (Reactant or reagent)
(prepn. of bicyclic heterocycles as humane peroxisome proliferator-activated receptor .gamma. agonists)
RN 213252-19-8 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 17 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



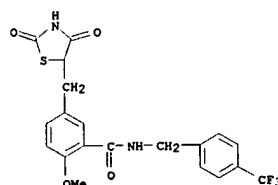
IT 223508-81-4P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP
(Preparation); RACT
(Reactant or reagent)
(prepn. of bicyclic heterocycles as humane peroxisome proliferator-activated receptor .gamma. agonists)
RN 223508-81-4 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-hydroxy-N-[(4-trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 18 OF 32 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:338335 CAPLUS
DOCUMENT NUMBER: 134:344604
TITLE: Antidiabetic formulation containing metformin and sulfonylurea
INVENTOR(S): Piper, Beth Anne
PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA
SOURCE: PCT Int. Appl., 76 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 2
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001032158	A2	20010510	WO 2000-US28467	20001013
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			US 1999-432465	A 19991103
PRIORITY APPLN. INFO.: AB A low dose antidiabetic formulation adapted for treating e.g., Type II diabetes contains a combination of metformin (at <800 mg/day) and at least 1 other antidiabetic agent such as a sulfonylurea. This combination provides at least about substantially equiv. efficacy in treating diabetes as do antidiabetic formulations contg. metformin employed in dosages prescribed in generally accepted medical practice for first line therapy in treating diabetes, but with substantially reduced side effects, such as hypoglycemia and/or gastrointestinal distress. A method for treating diabetes in drug naive human patients is also provided employing the above formulation to reduce insulin resistance and/or post-prandial glucose excursion and/or Hb 1AC, and/or increase post-prandial insulin, thereby				

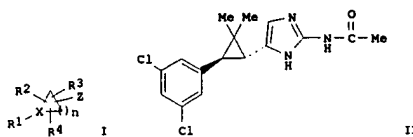
L4 ANSWER 18 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
treating the diabetes. Thus, tablets contained metformin-HCl 250.0, glyburide 1.25, croscarmellose sodium 7.00, Povidone 10.00, microcryst. cellulose 28.25, Mg stearate 2.25, and HPMC film-coating 6 mg. The effectiveness of this combination drug in producing hypoglycemia was demonstrated clin.
IT 213252-19-8, KRP-297
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(antidiabetic formulation contg. metformin and sulfonylurea)
RN 213252-19-8 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 19 OF 32 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:283949 CAPLUS
DOCUMENT NUMBER: 134:311218
TITLE: Synthesis and use of heterocyclic sodium/proton exchange inhibitors
INVENTOR(S): Ahmad, Saleem; Wu, Shung C.; O'Neill, Steven V.; Ngu, Khehyong; Atwal, Karnail S.
PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA
SOURCE: PCT Int. Appl., 221 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001027107	A2	20010419	WO 2000-US27461	20001002
WO 2001027107	A3	20020124		
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
EP 1224183	A2	20020724	EP 2000-968723	20001002
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL			
NO 2002001717	A	20020610	NO 2002-1717	20020411
PRIORITY APPLN. INFO.:			US 1999-158755P	P 19991012
			WO 2000-US27461	W 20001002
OTHER SOURCE(S):			MARPAT 134:311218	
GI				

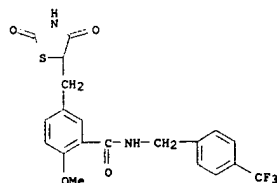
L4 ANSWER 19 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



AB Comps. of formula I [wherein: n is 1-5; X is N or CR5, where R5 is H, halo, alkenyl, alkynyl, alkoxy, alkyl, aryl or heteroaryl; Z is a heteroaryl group; R1 is H, alk(en)(yn)yl, alk(enyl)(ynyl)oxy, (aryl or alkyl)3Si, cycloalk(en)yl, (aryl)amino, aryl(alkyl), cycloheteroaryl, etc.; R2, R3 and R4 are any of the groups set out for R1 and optionally substituted with 1 to 5 substituents which may be the same or different and when X is N, R1 is preferably aryl or heteroaryl] are claimed. Several hundred examples are disclosed. Synthesis of II proceeds via cyclopropanation of the cinnamate derived from the olefination between 3,5-dichlorobenzaldehyde and t-butyl-diethylphosphonoacetate. The intermediate tert-Bu ester is converted to the corresponding .alpha.-chloroketone and reacted with acetyl guanidine to provide II in a total of 5 steps. Comps. I are said to be sodium/proton exchange inhibitors (NHE). Pharmaceutical combinations are claimed using I and certain antihypertensive agents, .beta.-adrenergic agonists, hypolipidemic agents, antidiabetic agents, antiobesity agents, etc. Comps. I are useful as antianginal and cardioprotective agents and provide a method for preventing or treating angina pectoris, cardiac dysfunction, myocardial necrosis, and arrhythmia. IT 213252-19-8, KRP297 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (pharmaceuticals also contg.; synthesis and use of heterocyclic sodium/proton exchange inhibitors)

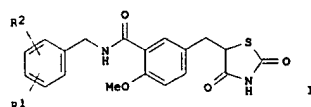
RN 213252-19-8 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 19 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



L4 ANSWER 20 OF 32 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:152661 CAPLUS
DOCUMENT NUMBER: 134:193428
TITLE: Preparation of substituted benzylthiazolidine-2,4-dione derivatives as agonists of human peroxisome proliferator-activated receptor
INVENTOR(S): Nomura, Masahiro; Murakami, Koji; Tsunoda, Masaki; Takahashi, Yukie
PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
SOURCE: PCT Int. Appl., 19 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001014352	A1	20010301	WO 2000-JP5522	20000818
W:	AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
EP 1207158	A1	20020522	EP 2000-953478	20000818
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL			
PRIORITY APPLN. INFO.:			JP 1999-235530	A 19990823
			WO 2000-JP5522	W 20000818
OTHER SOURCE(S):			MARPAT 134:193428	
GI				



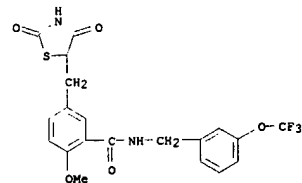
AB The title compds. (I), pharmaceutically acceptable salts thereof and hydrates of the same (wherein R1 represents chloro, bromo, nitro, trifluoromethoxy, ethoxy, propoxy or isopropoxy; and R2 represents hydrogen or chloro) are prepd. These compds. are capable of, as a ligand of human peroxisome proliferator-activated receptor (PPAR), enhancing the transcriptional activity of the receptor and showing effects of lowering blood sugar level and lowering lipid level; and a process for producing the same. Thus, 5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxybenzoic acid, Et3N, and CH2Cl2 were mixed, treated with Et chlorocarbonate and stirred under ice-cooling for 10 min, treated with 4-nitrobenzylamine, and then stirred at room temp. for 2 h to give 75% N-[(4-nitrophenyl)methyl]-5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxybenzamide (II). II and I (R1 = 4-n-Pro, R2 = H) enhanced the transcriptional activity of human PPAR.alpha. in CHO cells with EC50 of 0.53 and 0.11 .mu.M, resp.

IT 326926-46-9P 326926-47-0P 326926-48-1P
326926-49-2P 326926-50-5P 326926-51-6P
326926-52-7P 326926-53-8P 326926-54-9P,

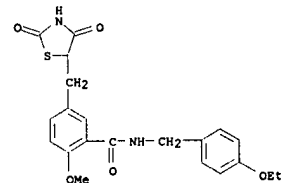
N-[(3,4-Dichlorophenyl)methyl]-5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxybenzamide
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (prepn. of substituted benzylthiazolidinedione deriva. as agonists of human peroxisome proliferator-activated receptor and blood sugar and lipid-lowering agents)

RN 326926-46-9 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-nitrophenyl)methyl]- (9CI) (CA INDEX NAME)

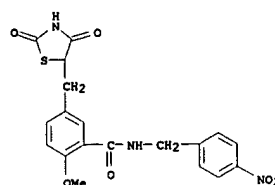
L4 ANSWER 20 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
RN 326926-49-2 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(3-trifluoromethoxyphenyl)methyl]- (9CI) (CA INDEX NAME)



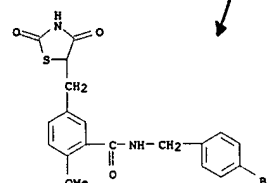
RN 326926-50-5 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-N-[(4-ethoxyphenyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



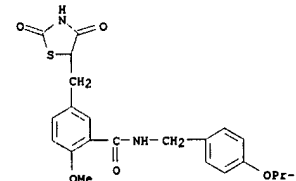
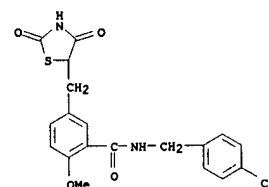
RN 326926-51-6 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-(1-methylethoxy)phenyl)methyl]- (9CI) (CA INDEX NAME)



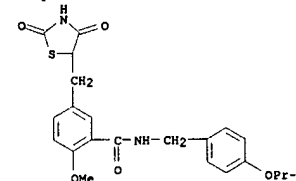
RN 326926-47-0 CAPLUS
CN Benzamide, N-[(4-bromophenyl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



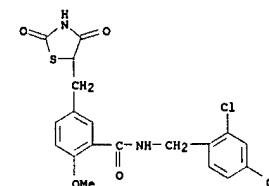
RN 326926-48-1 CAPLUS
CN Benzamide, N-[(4-chlorophenyl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



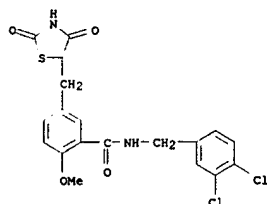
RN 326926-52-7 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-propoxyphenyl)methyl]- (9CI) (CA INDEX NAME)



RN 326926-53-8 CAPLUS
CN Benzamide, N-[(2,4-dichlorophenyl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



L4 ANSWER 20 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
RN 326926-54-9 CAPLUS
CN Benzamide, N-[(3,4-dichlorophenyl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



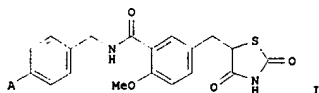
REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE
FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE
RE FORMAT

L4 ANSWER 21 OF 32 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:152660 CAPLUS
DOCUMENT NUMBER: 134:193427
TITLE: Preparation of substituted
benzylthiazolidine-2,4-
dione derivatives as agonists of human
peroxisome
proliferator-activated receptor
INVENTOR(S): Miyachi, Hiroyuki; Nomura, Masahiro; Tanase,
Takahiro;
PATENT ASSIGNEE(S): Murakami, Koji; Tsunoda, Masaki
Kyorin Pharmaceutical Co., Ltd., Japan
SOURCE: PCT Int. Appl., 20 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE
WO 2001014351 A1 20010301 WO 2000-05521 20000818
W: AE, AL, AM AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN,
CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG,
SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM,
AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE,
CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,
BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
EP 1207157 A1 20020522 EP 2000-953477 20000818
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE,
MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL
PRIORITY APPLN. INFO.: JP 1999-235529 A 19990823
JP 2000-242707 A 20000810
WO 2000-JP5521 W 20000818
OTHER SOURCE(S): MARPAT 134:193427
GI

3.01.01 NOT 371
g PCR
division?

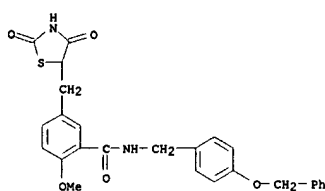
L4 ANSWER 21 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



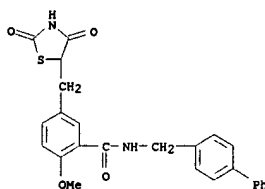
AB The title compds. represented by general formula (I; wherein A represents optionally substituted Ph, optionally substituted phenoxy or optionally substituted benzyloxy), pharmaceutically acceptable salts thereof and hydrates of the same are prepd. These compds. are capable of, as a ligand of human peroxisome proliferator-activated receptor (PPAR), enhancing the transcriptional activity of the receptor and showing effects of lowering blood sugar level and lowering lipid level. Thus, 5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxybenzoic acid, Et3N, and CH2Cl2 were mixed, treated with Et chlorocarbonate under ice-cooling, and stirred for 10 min under ice-cooling, followed by adding a soln. of 4-benzyloxybenzylamine in CH2Cl2, and the resulting mixt. was stirred at room temp. for 2 h to give 77% N-[(4-benzyloxyphenyl)methyl]-5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxybenzamide (II). II and I (A = PhO) enhanced the transcriptional activity of human PPAR.alpha. in CHO cells

with EC50 of 0.44 and 0.24 .mu.M, resp.
IT 326925-77-3P 326925-78-4P 326925-79-5P
326925-80-6P 326925-81-7P 326925-82-0P
326925-83-1P 326925-84-2P 326925-85-3P
326925-86-4P 326925-87-5P 326925-88-6P
326925-89-7P
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (prepn. of substituted benzylthiazolidinedione derivs. as agonists of human peroxisome proliferator-activated receptor and blood sugar and lipid-lowering agents)
RN 326925-77-3 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-phenylmethoxy)phenyl)methyl]- (9CI) (CA INDEX NAME)

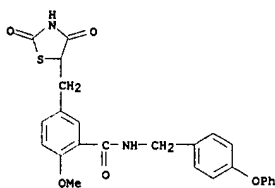
L4 ANSWER 21 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



RN 326925-78-4 CAPLUS
CN Benzamide, N-[(1,1'-biphenyl)-4-ylmethyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



RN 326925-79-5 CAPLUS
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-phenylmethoxy)phenyl)methyl]- (9CI) (CA INDEX NAME)

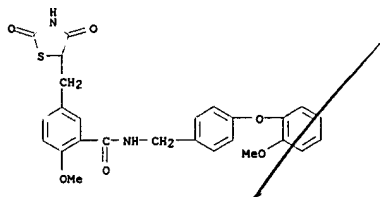


L4 ANSWER 21 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)

RN 326925-80-8 CAPLUS

CN Benzamide,

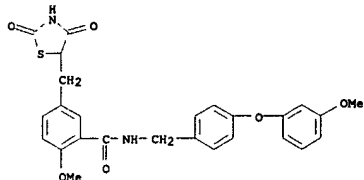
5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(2-methoxyphenoxy)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 326925-81-9 CAPLUS

CN Benzamide,

5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(3-methoxyphenoxy)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 326925-82-0 CAPLUS

CN Benzamide,

5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(4-methoxyphenoxy)phenyl]methyl]- (9CI) (CA INDEX NAME)

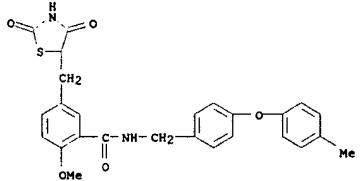


L4 ANSWER 21 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)

RN 326925-83-1 CAPLUS

CN Benzamide,

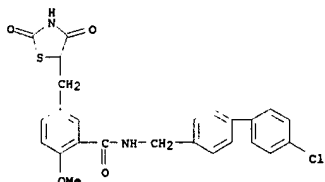
5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(3-methylphenoxy)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 326925-85-3 CAPLUS

CN Benzamide,

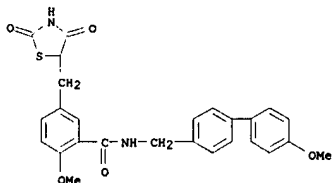
N-[[4'-(4-chlorophenyl)methoxy]biphenyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



RN 326925-86-4 CAPLUS

CN Benzamide,

5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4'-(4-methoxy[1,1'-biphenyl]-4-yl)methyl]- (9CI) (CA INDEX NAME)

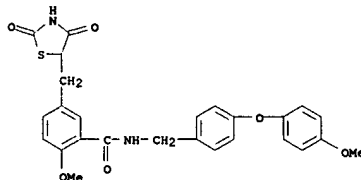


L4 ANSWER 21 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)

RN 326925-83-1 CAPLUS

CN Benzamide,

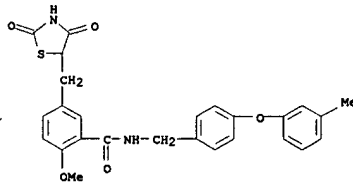
5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(3-methylphenoxy)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 326925-84-2 CAPLUS

CN Benzamide,

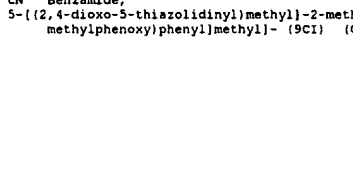
5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(4-methylphenoxy)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 326925-84-2 CAPLUS

CN Benzamide,

5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(4-methylphenoxy)phenyl]methyl]- (9CI) (CA INDEX NAME)

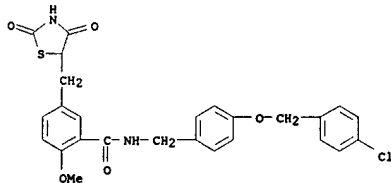


L4 ANSWER 21 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)

RN 326925-87-5 CAPLUS

CN Benzamide,

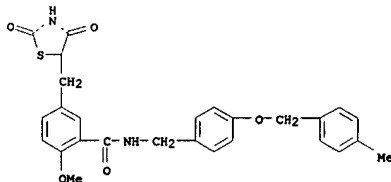
N-[[4'-(4-chlorophenyl)methoxy]biphenyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



RN 326925-88-6 CAPLUS

CN Benzamide,

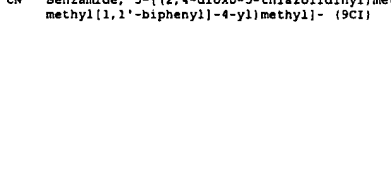
5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4'-(4-methylphenyl)methoxy]biphenyl]- (9CI) (CA INDEX NAME)

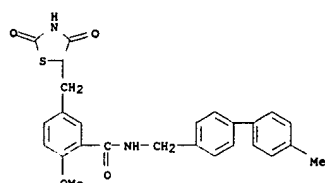


RN 326925-89-7 CAPLUS

CN Benzamide,

5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4'-(4-methyl[1,1'-biphenyl]-4-yl)methyl]- (9CI) (CA INDEX NAME)



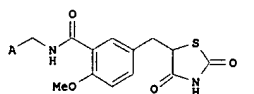


REFERENCE COUNT: 27 THERE ARE 27 CITED REFERENCES AVAILABLE
FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE
RE FORMAT

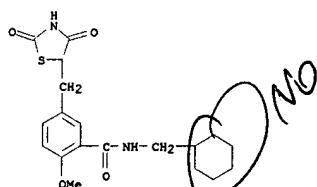
L4 ANSWER 22 OF 32 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2001:152659 CAPLUS
DOCUMENT NUMBER: 134:178551
TITLE: Preparation of substituted
benzylthiazolidine-2,4-
dione derivatives as ligands of human
peroxisome
proliferator-activated receptor
INVENTOR(S): Fujimori, Shizuyoshi; Murakami, Koji; Tsunoda,
Masaki
PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
SOURCE: PCT Int. Appl., 18 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE
WO 2001014350 A1 20010301 WO 2000-JP5520 20000818
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN,
CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG,
SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM,
AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE,
CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE,
BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
EP 1207156 A1 20020522 EP 2000-953476 20000818
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE,
MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL
PRIORITY APPLN. INFO.: JP 1999-235528 A 19990823
WO 2000-JP5520 W 20000818
GI

NO = NOT
371.



L4 ANSWER 22 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
AB The title compds. (I; wherein A represents pyridyl or cyclohexyl),
pharmaceutically acceptable salts thereof and hydrates of the same
are prepd. These compds. are capable of, as a ligand of human
peroxisome
proliferator-activated receptor (PPAR), enhancing the
transcriptional
activity of the receptor and showing effects of lowering blood
sugar level
and lowering lipid level. Thus,
5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-
methoxybenzoic acid, 2-picolylamine, 1-ethyl-3-(3-
dimethylaminopropyl)carbodiimide hydrochloride, and DMF were
stirred at
room temp. overnight to give 20% I (A = 2-pyridyl) (II). II and I
(A =
4-pyridyl) enhanced the transcriptional activity of human
PPAR.alpha. in
CHO cells with EC50 of 0.353 and 0.235 .mu.M, resp., and that of
human
PPAR.gamma. with EC50 of 0.30 and 0.14 .mu.M, resp.
IT 326922-18-3P
RI: BAC (Biological activity or effector, except adverse); BSU
(Biological
study, unclassified); SPN (Synthetic preparation); THU
(Therapeutic use);
BIOL (Biological study); PREP (Preparation); USES (Uses)
(prepn. of substituted benzylthiazolidinedione derivs. as
ligands of
human peroxisome proliferator-activated receptor and blood
sugar and
lipid-lowering agents)
RN 326922-18-3 CAPLUS
CN Benzamide,
N-(cyclohexylmethyl)-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-
methoxy- (9CI) (CA INDEX NAME)

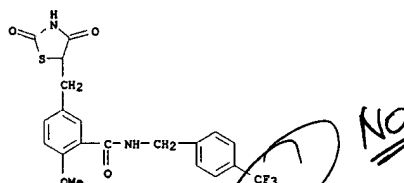


REFERENCE COUNT: 20 THERE ARE 20 CITED REFERENCES AVAILABLE
FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE
RE FORMAT

L4 ANSWER 23 OF 32 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2000:293502 CAPLUS
 DOCUMENT NUMBER: 133:84110
 TITLE: Fenofibrate and Rosiglitazone Lower Serum Triglycerides with Opposing Effects on Body Weight
 AUTHOR(S): Chaput, Evelyne; Saladin, Regis; Silvestre, Martine;
 CORPORATE SOURCE: Edgar, Alan D.
 Department of Metabolic Diseases, Laboratoire Fournier, Daix, 21121, Fr.
 SOURCE: Biochemical and Biophysical Research Communications
 (2000), 271(2), 445-450
 CODEN: BBRCA9; ISSN: 0006-291X
 PUBLISHER: Academic Press
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Activators of peroxisome proliferator activated receptors (PPARs) are effective drugs to improve the metabolic abnormalities linking hypertriglyceridemia to diabetes, hyperglycemia, insulin-resistance, and atherosclerosis. We compared the pharmacol. profile of a PPAR.alpha. activator, fenofibrate, and a PPAR.gamma. activator, rosiglitazone, on serum parameters, target gene expression, and body wt. gain in (fa/fa) fatty Zucker rats and db/db mice as well as their assocn. in db/db mice. Fenofibrate faithfully modified the expression of PPAR.alpha. responsive genes. Rosiglitazone increased adipose tissue aP2 mRNA in both models while increasing liver acyl CoA oxidase mRNA in db/db mice but not in fatty Zucker rats. Both drugs lowered serum triglycerides yet rosiglitazone markedly increased body wt. gain while fenofibrate decreased body wt. gain in fatty Zucker rats. KRP 297, which has been reported to be a PPAR.alpha. and .gamma. co-activator, also affected serum triglycerides and insulin in fatty Zucker rats although no change in body wt. gain was noted. These results serve to clearly differentiate the metabolic finality of two distinct classes of drugs, as well as their corresponding nuclear receptors, having similar effects on serum triglycerides. (c) 2000 Academic Press.
 IT 213252-19-8, KRP 297
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)
 (fenofibrate and rosiglitazone lower serum triglycerides with opposing

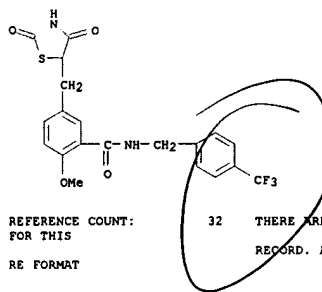
L4 ANSWER 24 OF 32 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2000:243901 CAPLUS
 DOCUMENT NUMBER: 133:12622
 TITLE: Tissue-specific actions of antidiabetic thiazolidinediones on the reduced fatty acid oxidation in skeletal muscle and liver of Zucker diabetic fatty rats
 AUTHOR(S): Ide, Tomohiro; Nakazawa, Tomoko; Mochizuki, Toshiro;
 CORPORATE SOURCE: Murakami, Koji
 Central Research Laboratories, Kyorin Pharmaceutical,
 Tochigi, 329-0114, Japan
 SOURCE: Metabolism, Clinical and Experimental (2000), 49(4), 521-525
 CODEN: METAJ; ISSN: 0026-0495
 PUBLISHER: W. B. Saunders Co.
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB Fatty acid overload has been proposed as a cause of decreased responsiveness in the major insulin target tissues of the body such as muscle and liver tissue. We therefore investigated fatty acid oxidn. in soleus muscle and liver isolated from Zucker diabetic fatty (ZDF) rats treated with thiazolidinediones, a new class of antidiabetic agents. 14CO2 prodn. from [14C]palmitic (C16:0) acid was lower in the soleus muscle and liver of ZDF rats vs. lean rats (P < .05). When administered orally to ZDF rats for 2 wk, the thiazolidinediones troglitazone (300 mg/kg) and KRP-297 (10 mg/kg) increased palmitic acid oxidn. in the soleus muscle of ZDF rats (P < .05). KRP-297, but not troglitazone, increased palmitic acid oxidn. in the liver of ZDF rats (P < .05), and both troglitazone and KRP-297 inhibited triglyceride accumulation in the skeletal muscle of ZDF rats. Hepatic triglyceride accumulation in ZDF rats was inhibited by KRP-297, but not by troglitazone. A redn. of fatty acid oxidn. in the liver of ZDF rats and an increase in response to KRP-297 were obsd. only when C16:0 and C18:0 fatty acids, not C8:0, were used as substrates. Thus, there were defects in fatty acid catabolic activity and triglyceride accumulation in the soleus muscle and liver of ZDF rats. These results indicate that KRP-297 has advantages over troglitazone in the amelioration of these lipid metabolic abnormalities in

L4 ANSWER 23 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
 effects on body wt.)
 RN 213252-19-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 2 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 24 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
 insulin resistance assocd. with obesity.
 IT 213252-19-8, KRP-297
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (tissue-specific actions of antidiabetic thiazolidinediones on reduced fatty acid oxidn. in muscle and liver in NIDDM/obesity)
 RN 213252-19-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

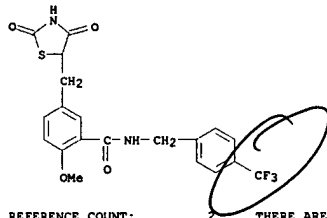
L4 ANSWER 25 OF 32 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 2000:190928 CAPLUS
 DOCUMENT NUMBER: 132:231969
 TITLE: Method for treating diabetes employing an α P2 inhibitor and combination
 INVENTOR(S): Robl, Jeffrey A.; Parker, Rex A.; Biller, Scott A.;
 Krishna
 PATENT ASSIGNEE(S): Bristol-Myers Squibb Co., USA
 SOURCE: PCT Int. Appl., 55 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000015229	A1	20000323	WO 1999-US20946	19990913
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
AU 9963877	A1	20000403	AU 1999-63877	19990913
BR 9913833	A	20010529	BR 1999-13833	19990913
EP 1121129	A1	20010808	EP 1999-951438	19990913
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
NO 2001001351	A	20010511	NO 2001-1351	20010316
LT 4871	B	20011227	LT 2001-22	20010316
LT 4870	B	20011227	LT 2001-23	20010316
US 2002035064	A1	20020321	US 2001-905235	20010713

PRIORITY APPLN. INFO.:
 US 1998-100677P P 19980917
 US 1999-390275 B1 19990907
 WO 1999-US20946 W 19990913
 OTHER SOURCE(S): MARPAT 132:231969
 AB A method is provided for treating diabetes and related diseases, such as insulin resistance, obesity, hyperglycemia, hyperinsulinemia, elevated blood levels of free fatty acids or glycerol, hypertriglyceridemia, and esp. Type II diabetes, employing an adipocyte protein α P2 inhibitor or a

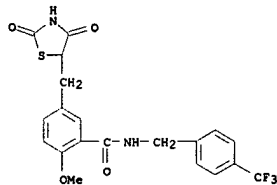
L4 ANSWER 26 OF 32 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 1999:751167 CAPLUS
 DOCUMENT NUMBER: 132:44794
 TITLE: Amelioration by KRP-297, a new thiazolidinedione, of impaired glucose uptake in skeletal muscle from obese insulin-resistant animals
 AUTHOR(S): Murakami, Koji; Tsunoda, Masaki; Ide, Tomohiro; Ohashi, Mitsuo; Mochizuki, Toshiro
 CORPORATE SOURCE: Central Research Laboratories, Kyorin Co Ltd., Tochigi, Japan
 SOURCE: Metabolism, Clinical and Experimental (1999), 48(11), 1450-1454
 CODEN: METAJ; ISSN: 0026-0495
 PUBLISHER: W. B. Saunders Co.
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB We examd. the effect of KRP-297, a new thiazolidinedione deriv., on glucose uptake in the soleus muscle of two animal models of insulin resistance that show moderate (ob/ob mice) and severe (db/db mice) hyperglycemia. Insulin-stimulated 2-deoxyglucose (2DG) uptake in soleus muscle was 53.8% lower in ob/ob mice vs. lean mice ($P < .05$). When administered to ob/ob mice, KRP-297 (0.3 to 10 mg/kg) decreased plasma glucose and insulin levels and improved the impaired insulin-stimulated 2DG uptake in soleus muscle in a dose-dependent manner. Soleus muscle from db/db mice exhibited defects in both basal (35.0% decrease, $P < .01$) and insulin-stimulated (50.5% decrease, $P < .01$) 2DG uptake. These defects were improved by treatment with KRP-297 (0.3 to 10 mg/kg). Moreover, KRP-297 prevented severe hyperglycemia and the marked decrease in pancreatic insulin content in db/db mice. These results suggest that KRP-297 treatment is useful to prevent the development of diabetic syndromes in addn. to ameliorating the impaired glucose transport in skeletal muscle.
 IT 213252-19-8, KRP-297
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (thiazolidinedione deriv. KRP-297 amelioration of impaired glucose uptake in skeletal muscle from obese insulin-resistant animals)
 RN 213252-19-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 25 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
 combination of an α P2 inhibitor and another antidiabetic agent such as metformin, glyburide, troglitazone and/or insulin.
 IT 213252-19-8, KRP 297
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (α P2 inhibitor and combination with another antidiabetic agent for treatment of diabetes and related diseases)
 RN 213252-19-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

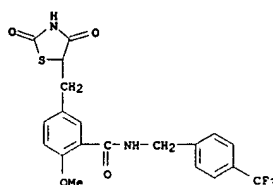
L4 ANSWER 26 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



REFERENCE COUNT: 23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 27 OF 32 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 1999:436161 CAPLUS
 DOCUMENT NUMBER: 131:238315
 TITLE: Evidence for direct binding of fatty acids and eicosanoids to human peroxisome proliferator-activated receptor .alpha.
 AUTHOR(S): Murakami, Koji; Ide, Tomohiro; Suzuki, Masahiro;
 CORPORATE SOURCE: Mochizuki, Toshiro; Kadowaki, Takashi; Central Research Laboratories, Kyorin Co., Ltd., Tochigi, Japan
 SOURCE: Biochemical and Biophysical Research Communications
 (1999), 260(3), 609-613
 CODEN: BBRC99; ISSN: 0006-291X
 PUBLISHER: Academic Press
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB The .alpha. isoform of peroxisome proliferator-activated receptor (PPAR) is activated by fatty acids, their metabolites, and the fibrates class of lipid-lowering agents. To test the ability of these activators to directly bind the ligand-binding domain of human PPAR.alpha., we performed a competitive binding assay using radiolabeled [3H]KRP-297, a known ligand for human PPAR.alpha.. Long-chain fatty acids and eicosanoids were even more potent ligands for human PPAR.alpha. than the hitherto most potent PPAR.alpha. ligand WY-14,643. Moreover, these natural ligands avidly activated this receptor in a transient transcriptional assay. This study provides the direct evidence that human PPAR.alpha. is activated through the direct binding of fatty acids and eicosanoids, as well as of a fibrates, to its ligand-binding domain. (c) 1999 Academic Press.
 IT 213252-19-8, KRP-297
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)
 (direct binding of fatty acids and eicosanoids to human peroxisome proliferator-activated receptor .alpha.)
 RN 213252-19-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 27 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)

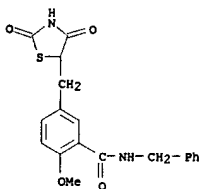


REFERENCE COUNT: 24 THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

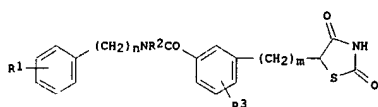
2/22/99 = good date

L4 ANSWER 28 OF 32 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 1999:188591 CAPLUS
 DOCUMENT NUMBER: 130:311725
 TITLE: (3-Substituted benzyl)thiazolidine-2,4-diones as structurally new antihyperglycemic agents
 AUTHOR(S): Nomura, Masahiro; Kinoshita, Susumu; Satoh, Naeda, Toshio; Murakami, Koji; Tsunoda, Masaki; Miyachi, Hiroyuki; Awano, Katsuya
 CORPORATE SOURCE: Central Research Laboratories, Kyorin Co., Ltd., Tochigi, 329-0114, Japan
 SOURCE: Bioorganic & Medicinal Chemistry Letters
 (1999), 9(4), 533-538
 CODEN: BMCLE8; ISSN: 0960-894X
 PUBLISHER: Elsevier Science Ltd.
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 GI

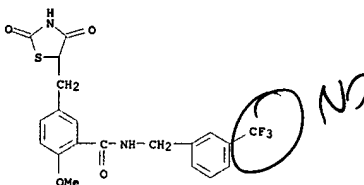
L4 ANSWER 28 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



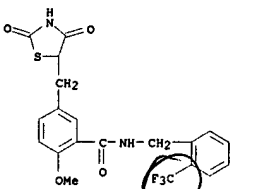
RN 185808-40-6 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(3-(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)



AB Title compds. I (R1 = 4-tert-Bu, H, 4-Me, 4-MeO, 4-CF3, etc.; R2 = H, Et; R3 = 6-MeO, 4-MeO, 2-MeO, 6-EtO, 6-OH, 6-F, etc.; m = 0-3; n = 0-2) were prepd. A structure-activity study of these compds. led to the identification of I (R1 = CF3, R2 = H, R3 = 6-MeO, m = n = 1) (KRP-297) as a candidate drug for the treatment of diabetes mellitus.
 IT 185808-38-2P 185808-40-6P 185808-42-8P 185808-45-1P 185808-49-5P 185808-51-9P 185808-62-2P 185808-63-3P 185808-64-4P 185808-65-5P 185808-67-7P 185808-68-8P 185808-70-2P 186312-86-7P 213252-19-8P, KRP-297 223508-81-4P
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)
 (prepn. and antihyperglycemic activity of)
 RN 185808-38-2 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-(phenylmethyl)- (9CI) (CA INDEX NAME)

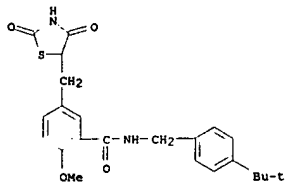


RN 185808-42-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(2-(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)

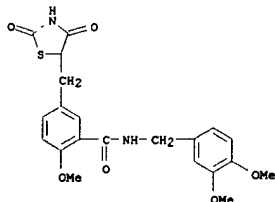


RN 185808-45-1 CAPLUS

L4 ANSWER 28 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
 CN Benzamide, N-[[[4-(1,1-dimethylethyl)phenyl]methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)

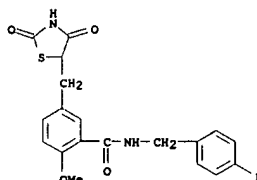


RN 185808-49-5 CAPLUS
 CN Benzamide, N-[[[3,4-dimethoxyphenyl]methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)

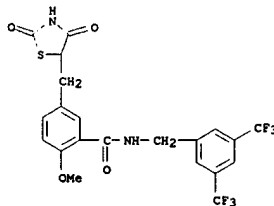


RN 185808-51-9 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-N-[(4-fluorophenyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)

L4 ANSWER 28 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)

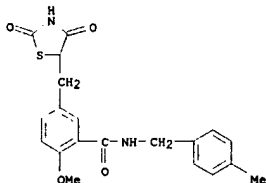


RN 185808-62-2 CAPLUS
 CN Benzamide, N-[[[3,5-bis(trifluoromethyl)phenyl]methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)

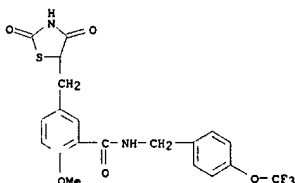


RN 185808-63-3 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-methylphenyl)methyl]- (9CI) (CA INDEX NAME)

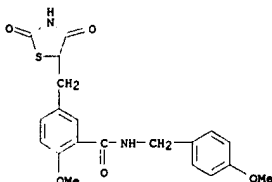
L4 ANSWER 28 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



RN 185808-64-4 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-(trifluoromethoxy)phenyl)methyl]- (9CI) (CA INDEX NAME)

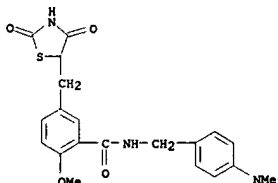


RN 185808-65-5 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-methoxyphenyl)methyl]- (9CI) (CA INDEX NAME)

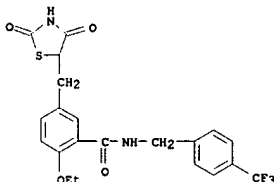


L4 ANSWER 28 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)

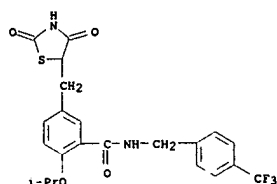
RN 185808-67-7 CAPLUS
 CN Benzamide, N-[[[4-(dimethylamino)phenyl]methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



RN 185808-68-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-ethoxy-N-[(4-(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)



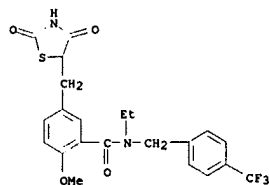
RN 185808-70-2 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-(1-methylethoxy)-N-[(4-(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)



RN 186312-86-7 CAPLUS

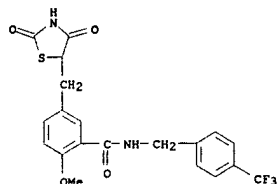
CN Benzamide,

5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 29 OF 32 CAPLUS COPYRIGHT 2002 ACS

ACCESSION NUMBER: 1998:784882 CAPLUS

DOCUMENT NUMBER: 130:148506

TITLE: A novel insulin sensitizer acts as a coligand for peroxisome proliferator-activated

receptor-.alpha.

metabolism

AUTHOR(S): Murakami, Koji; Tobe, Kazuyuki; Ide, Tomohiro; Mochizuki, Toshiro; Ohashi, Mitsuo; Akanuma,

Yasuo;

CORPORATE SOURCE: Yazaki, Yoshio; Kadowaki, Takashi
Third Department of Internal Medicine, Faculty
of Medicine, University of Tokyo, Tokyo, 113,

Japan

SOURCE: Diabetes (1998), 47(12), 1841-1847

CODEN: DIAEAS; ISSN: 0012-1797

PUBLISHER: American Diabetes Association

DOCUMENT TYPE: Journal

LANGUAGE: English

AB We investigated the biol. activity of a novel thiazolidinedione

(TZD)

deriv., KRP-297, and the mol. basis of this activity. When

administered

to obese Zucker fatty rats (obese rats) at 10 mg/kg for 2 wk,

KRP-297,

unlike BRL-49653, restored reduced lipid oxidn., i.e., CO2 and

ketone body

prodn. from [14C]palmitic acid, in the liver by 39% (P < 0.05) and

57% (P

< 0.01), resp. KRP-297 was also significantly more effective than

BRL-49653 in the inhibition of enhanced lipogenesis and

triglyceride

accumulation in the liver. To understand the mol. basis of the

biol.

effects of KRP-297, we examd. the effect on peroxisome

proliferator-activated receptor (PPAR) isoforms, which may play

key roles

in lipid metab. Unlike classical TZD derivs., KRP-297 activated

both

PPAR-.alpha. and PPAR-.gamma., with median effective concns. of

1.0 and

0.8 .mu.mol/L, resp. Moreover, radiolabeled [3H]KRP-297 bound

directly to

PPAR-.alpha. and PPAR-.gamma. with dissocn. consts. of 228 and 326

nmol/L,

resp. Concomitantly, KRP-297, but not BRL-49653, increased the

mRNA

and the activity (1.5-fold [P < 0.01] and 1.8-fold [P < 0.05], resp.)

of

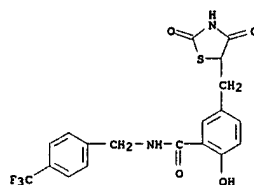
acyl-CoA oxidase, which has been reported to be regulated by

PPAR-.alpha.,

in the liver. By contrast, KRP-297 (P < 0.05) was less potent than

RN 223508-81-4 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-hydroxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)

REFERENCE COUNT:
FOR THIS

15 THERE ARE 15 CITED REFERENCES AVAILABLE

RE FORMAT

RECORD. ALL CITATIONS AVAILABLE IN THE

L4 ANSWER 29 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)

BRL-49653 (P < 0.01) in inducing the PPAR-.gamma.-regulated ap2

gene mRNA

expression in the adipose tissues. These results suggest that

PPAR-.alpha. agonism has a protective effect against abnormal

lipid metab.

in liver of obese rats.

IT 213252-19-8, KRP 297

RL: BAC (Biological activity or effector, except adverse); BSU

(Biological

study, unclassified); THU (Therapeutic use); BIOL (Biological

study); USES

(Uses)

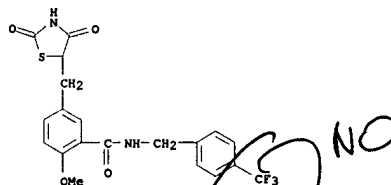
(effect of PPAR-.alpha. activation by insulin sensitizer,

thiazolidinedione deriv. KRP-297, on abnormal lipid metab. in

liver of Zucker fatty rats)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)

REFERENCE COUNT:
FOR THIS

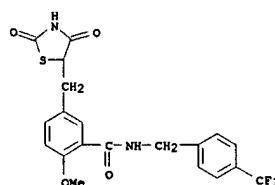
35 THERE ARE 35 CITED REFERENCES AVAILABLE

RE FORMAT

RECORD. ALL CITATIONS AVAILABLE IN THE

L4 ANSWER 30 OF 32 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 1998:421607 CAPLUS
 DOCUMENT NUMBER: 129:239719
 TITLE: Effects of PPAR.alpha. activation on liver lipid metabolism in Zucker fatty rats
 AUTHOR(S): Ide, Tomohiro; Murakami, Koji; Tobe, Kazuyuki; Mochizuki, Toshiro; Ohashi, Mitsuo; Akanuma, Yasuo;
 Kadowaki, Takashi; Yazaki, Yoshio
 CORPORATE SOURCE: Cent. Res. Lab., Kyorin Pharm. Co., Ltd., Tochigi,
 329-01, Japan
 SOURCE: Diabetes Frontier (1998), 9(3), 345-346
 CODEN: DIFREZ; ISSN: 0915-6593
 PUBLISHER: Medikaru Rebyusha
 DOCUMENT TYPE: Journal
 LANGUAGE: Japanese
 AB Oral administration of KRP-297 or BRL-49653 with high affinity to PPAR .alpha. to Zucker fatty (obese) rats and to control lean rats for 2 wk significantly lowered the blood glucose, insulin, triglyceride, and free fatty acid levels in the obese rats. KRP-297 and BRL-49653 also suppressed the increase in triglyceride accumulation and fatty acid biosynthesis activity in the liver of the obese rats as compared to the lean rats. In contrast, the markedly reduced activity of the hepatic acyl-CoA oxidase in the obese rats was markedly recovered by the administration. The results suggest that the activation of PPAR .alpha. by KRP-297 or BRL-49653 (ligand) might have inhibitory action on the hepatic triglyceride accumulation and lipid metab. abnormality in the obese rats.
 IT 213252-19-8, KRP 297
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study) (effects of PPAR.alpha. activation on liver lipid metab. in Zucker fatty rats)
 RN 213252-19-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)

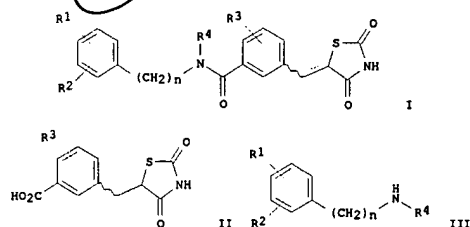
L4 ANSWER 30 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



L4 ANSWER 31 OF 32 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 1997:116453 CAPLUS
 DOCUMENT NUMBER: 126:157499
 TITLE: Preparation of N-substituted dioxothiazolidylbenzamide derivatives as blood sugar lowering agents
 INVENTOR(S): Maeda, Toshio; Nomura, Masahiro; Awano, Katsuya;
 Kinoshta, Susumu; Sato, Hiroya; Murakami, Koji;
 Tsunoda, Masaki
 PATENT ASSIGNEE(S): Kyorin Seiyaku Kk, Japan
 SOURCE: Jpn. Kokai Tokkyo Koho, 11 pp. CODEN: JKKXAF
 Patent
 DOCUMENT TYPE: Japanese
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

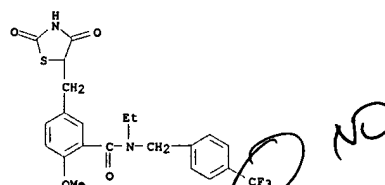
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 5333355	A2	19961217	JP 1995-159782	19950602

OTHER SOURCE(S): MARPAT 126:157499
 GI



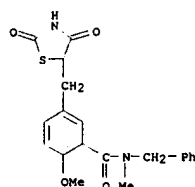
AB The title compds. (I; R1, R2 = H, Cl-4 alkyl, Cl-3 alkoxy, haloalkoxy, or haloalkyl, halo, OH, NO2, etc.; R3 = H, Cl-3 alkoxy, halo, OH; R4 = H, Cl-4 alkyl; dotted line = single or double bond; n = 0-2) are prepd. by reacting benzoic acid derivs. (II; R3, dotted line = same as above) with amines (III; R1, R2, R4, n = same as above). I, possessing blood sugar and lipid lowering activities, are useful for diabetes mellitus and hyperlipemia. Thus, 5-(2,4-dioxothiazolidyl-5-ylidene)methyl-2-methoxybenzoic acid was reacted with 4-tert-butylaniline in the presence

L4 ANSWER 31 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
 of Et3N and NCP(O)(OEt)2 to give 99% I (R1 = 4-tert-BuC6H4, R3 = 2-MeO, R2 = R4 = H, dotted line = double bond, n = 0). I (R1 = R2 = 4-CF3, R3 = 6-MeO, R4 = Et, dotted line = single bond, n = 1) at 10 mg/kg showed 31% blood sugar lowering activity when tested on mice p.o. in vivo.
 IT 186312-86-7P 186312-87-8P
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (prepn. of N-substituted dioxothiazolidylbenzamide derivs. as blood sugar lowering agents)
 RN 186312-86-7 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-N-ethyl-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 186312-87-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-methyl-N-(phenylmethyl)- (9CI) (CA INDEX NAME)

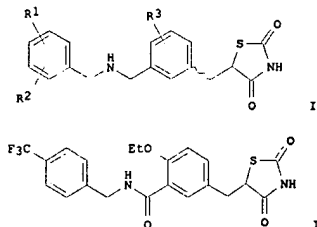
10% generic
 teach
 genus.



ACCESSION NUMBER: 1997:85180 CAPLUS
 DOCUMENT NUMBER: 126:104076
 TITLE: Preparation of derivatives as antidiabetics and hypolipemics
 INVENTOR(S): Maeda, Toshio; Nomura, Masahiro; Awano, Katsuya;
 Katsuya;
 Kinoshita, Susumu; Satoh, Hiroya; Murakami, Koji;
 Tsunoda, Masaki
 PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan; Maeda, Toshio;
 Nomura, Masahiro; Awano, Katsuya; Kinoshita, Susumu;
 Satoh, Hiroya; Murakami, Koji; Tsunoda, Masaki
 SOURCE: PCT Int. Appl., 40 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

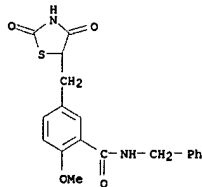
12.5.96

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9638428	A1	19961205	WO 1996-JP1459	19960530
W: AU, CA, CN, HU, KR, US				
RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
JP 09048771	A2	19970218	JP 1996-153139	19960524
JP 3144624	B2	20010312		
JP 2001139565	A2	20010522	JP 2000-350367	19960524
CA 2220698	AA	19961205	CA 1996-2220698	19960530
AU 9658446	A1	19961218	AU 1996-58446	19960530
AU 698896	B2	19981112		
EP 846693	A1	19980610	EP 1996-920002	19960530
EP 846693	B1	20020123		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, PT, IE, FI				
CN 1186489	A	19980701	CN 1996-194390	19960530
CN 1069901	B	20010822		
AT 212341	E	20020215	AT 1996-920002	19960530
TW 400328	B	20000801	TW 1996-85106555	19960601
US 6030990	A	20000229	US 1997-952672	19971202
US 6001862	A	19991214	US 1999-292955	19990416
US 6147101	A	20001114	US 2000-482268	20000113
CN 1336366	A	20020220	CN 2000-130138	20001017
PRIORITY APPL. INFO.:			JP 1995-159781	A 19950602
			JP 1996-153139	A 19960524
			WO 1996-JP1459	W 19960530
OTHER SOURCE(S):			MARPAT 126:104076	
GI				

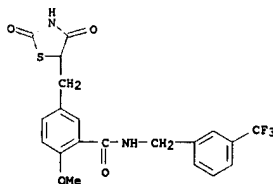


AB Novel N-benzylthiazolidinylbenzamide derivs. represented by general formula I (R1 and R2 are the same or different and each represents hydrogen, lower (C1-4) alkyl, lower (C1-3) alkoxy, lower (C1-3) haloalkyl, lower (C1-3) haloalkoxy, halogeno, hydroxy, nitro, amino optionally substituted by lower (C1-3) alkyl or a heterocycle, or R1 and R2 may be bonded to each other to form methylenedioxy; R3 represents lower (C1-3) alkoxy, hydroxy or halogeno; and the dotted line represents a double or single bond] are prepd. The title compd. II at 10 mg/kg gave 37% decrease in blood sugar in obese mice.

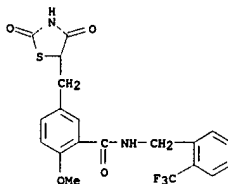
IT 185808-38-2P 185808-40-6P 185808-42-8P
 185808-45-1P 185808-49-5P 185808-51-9P
 185808-62-2P 185808-63-3P 185808-64-4P
 185808-65-5P 185808-66-6P 185808-67-7P
 185808-68-8P 185808-69-9P 185808-70-2P
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (prepn. of N-benzylthiazolidinylbenzamide derivs. as antidiabetics and hypolipemics)
 RN 185808-38-2 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(phenylmethyl)- (9CI) (CA INDEX NAME)]



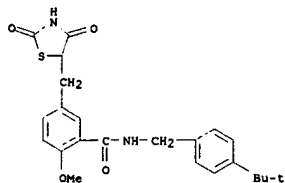
RN 185808-40-6 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(3-(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)



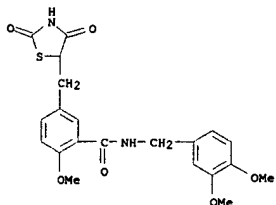
RN 185808-42-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(2-(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 32 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)
 RN 185808-45-1 CAPLUS
 CN Benzamide, N-[(4-{(1,1-dimethylethyl)phenyl)methyl}-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)

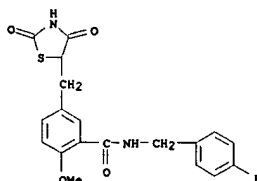


RN 185808-49-5 CAPLUS
 CN Benzamide, N-[(3,4-dimethoxyphenyl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)

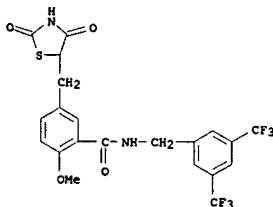


RN 185808-51-9 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-N-[(4-fluorophenyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)

L4 ANSWER 32 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)

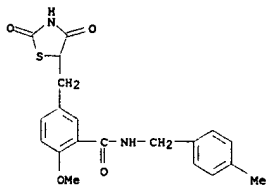


RN 185808-62-2 CAPLUS
 CN Benzamide, N-[(3,5-bis(trifluoromethyl)phenyl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)

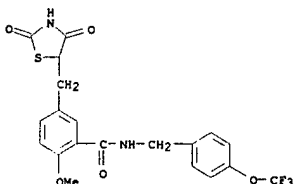


RN 185808-63-3 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-methylphenyl)methyl]- (9CI) (CA INDEX NAME)

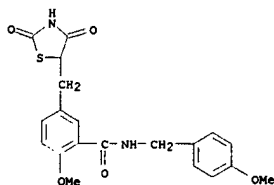
L4 ANSWER 32 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)



RN 185808-64-4 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-(trifluoromethoxy)phenyl)methyl]- (9CI) (CA INDEX NAME)

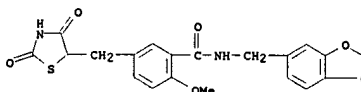


RN 185808-65-5 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-methoxyphenyl)methyl]- (9CI) (CA INDEX NAME)

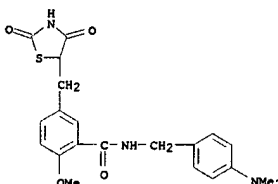


L4 ANSWER 32 OF 32 CAPLUS COPYRIGHT 2002 ACS (Continued)

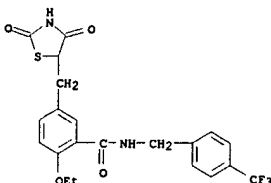
RN 185808-66-6 CAPLUS
 CN Benzamide, N-[(1,3-benzodioxol-5-yl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



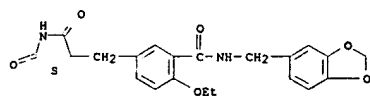
RN 185808-67-7 CAPLUS
 CN Benzamide, N-[(4-(dimethylamino)phenyl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



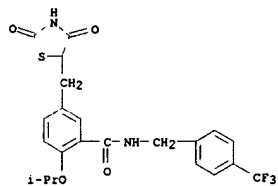
RN 185808-68-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-ethoxy-N-[(4-(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)



RN 185808-69-9 CAPLUS
 CN Benzamide, N-[(1,3-benzodioxol-5-yl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-ethoxy- (9CI) (CA INDEX NAME)



RN 185808-70-2 CAPLUS
 CN Benzamide,
 5-({(2,4-dioxo-5-thiazolidinyl)methyl}-2-(1-methylethoxy)-N-({(4-(trifluoromethyl)phenyl)methyl}- (9CI) (CA INDEX NAME)



10049937

Connecting via Winsock to STN

LOGINID:
SSSPTA1613SXW

STNLOGON timed out

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID:SSSPTA1613SXW

PASSWORD:
TERMINAL (ENTER 1, 2, 3, OR ?):2

* * * * * Welcome to STN International * * * * *

NEWS	1		Web Page URLs for STN Seminar Schedule - N. America
NEWS	2	Apr 08	"Ask CAS" for self-help around the clock
NEWS	3	Apr 09	BEILSTEIN: Reload and Implementation of a New Subject Area
NEWS	4	Apr 09	ZDB will be removed from STN
NEWS	5	Apr 19	US Patent Applications available in IFICDB, IFIPAT, and IFIUDB
NEWS	6	Apr 22	Records from IP.com available in CAPLUS, HCAPLUS, and ZCAPLUS
NEWS	7	Apr 22	BIOSIS Gene Names now available in TOXCENTER
NEWS	8	Apr 22	Federal Research in Progress (FEDRIP) now available
NEWS	9	Jun 03	New e-mail delivery for search results now available
NEWS	10	Jun 10	MEDLINE Reload
NEWS	11	Jun 10	PCTFULL has been reloaded
NEWS	12	Jul 02	FOREGE no longer contains STANDARDS file segment
NEWS	13	Jul 22	USAN to be reloaded July 28, 2002; saved answer sets no longer valid
NEWS	14	Jul 29	Enhanced polymer searching in REGISTRY
NEWS	15	Jul 30	NETFIRST to be removed from STN
NEWS	16	Aug 08	CANCERLIT reload
NEWS	17	Aug 08	PHARMAMarketLetter(PHARMAML) - new on STN
NEWS	18	Aug 08	NTIS has been reloaded and enhanced
NEWS	19	Aug 19	Aquatic Toxicity Information Retrieval (AQUIRE) now available on STN
NEWS	20	Aug 19	IFIPAT, IFICDB, and IFIUDB have been reloaded
NEWS	21	Aug 19	The MEDLINE file segment of TOXCENTER has been reloaded
NEWS	22	Aug 26	Sequence searching in REGISTRY enhanced
NEWS	23	Sep 03	JAPIO has been reloaded and enhanced
NEWS	24	Sep 16	Experimental properties added to the REGISTRY file
NEWS	25	Sep 16	CA Section Thesaurus available in CAPLUS and CA
NEWS	26	Oct 01	CASREACT Enriched with Reactions from 1907 to 1985
NEWS	27	Oct 21	EVENTLINE has been reloaded
NEWS	28	Oct 24	BEILSTEIN adds new search fields
NEWS	29	Oct 24	Nutraceuticals International (NUTRACEUT) now available on STN
NEWS	30	Oct 25	MEDLINE SDI run of October 8, 2002
NEWS	31	Nov 18	DKILIT has been renamed APOLLIT
NEWS	32	Nov 25	More calculated properties added to REGISTRY
NEWS	33	Dec 02	TIBKAT will be removed from STN

10049937

NEWS 34 Dec 04 CSA files on STN
NEWS 35 Dec 17 PCTFULL now covers WP/PCT Applications from 1978 to date
NEWS 36 Dec 17 TOXCENTER enhanced with additional content
NEWS 37 Dec 17 Adis Clinical Trials Insight now available on STN
NEWS 38 Dec 30 ISMEC no longer available
NEWS 39 Jan 21 NUTRACEUT offering one free connect hour in February 2003
NEWS 40 Jan 21 PHARMAML offering one free connect hour in February 2003
NEWS 41 Jan 29 Simultaneous left and right truncation added to COMPENDEX,
ENERGY, INSPEC
NEWS 42 Feb 13 CANCERLIT is no longer being updated
NEWS 43 Feb 24 METADEX enhancements
NEWS 44 Feb 24 PCTGEN now available on STN
NEWS 45 Feb 24 TEMA now available on STN
NEWS 46 Feb 26 NTIS now allows simultaneous left and right truncation
NEWS 47 Feb 26 PCTFULL now contains images
NEWS 48 Mar 04 SDI PACKAGE for monthly delivery of multifile SDI results
NEWS 49 Mar 19 APOLLIT offering free connect time in April 2003
NEWS 50 Mar 20 EVENTLINE will be removed from STN
NEWS 51 Mar 24 PATDPAFULL now available on STN
NEWS 52 Mar 24 Additional information for trade-named substances without
structures available in REGISTRY
NEWS 53 Mar 24 Indexing from 1957 to 1966 added to records in CA/CAPLUS

NEWS EXPRESS January 6 CURRENT WINDOWS VERSION IS V6.01a,
CURRENT MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP),
AND CURRENT DISCOVER FILE IS DATED 01 OCTOBER 2002
NEWS HOURS STN Operating Hours Plus Help Desk Availability
NEWS INTER General Internet Information
NEWS LOGIN Welcome Banner and News Items
NEWS PHONE Direct Dial and Telecommunication Network Access to STN
NEWS WWW CAS World Wide Web Site (general information)

Enter NEWS followed by the item number or name to see news on that
specific topic.

All use of STN is subject to the provisions of the STN Customer
agreement. Please note that this agreement limits use to scientific
research. Use for software development or design or implementation
of commercial gateways or other similar uses is prohibited and may
result in loss of user privileges and other penalties.

* * * * * STN Columbus * * * * *

FILE 'HOME' ENTERED AT 08:57:01 ON 26 MAR 2003

=> FIL REG

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	0.21	0.21

FILE 'REGISTRY' ENTERED AT 08:57:10 ON 26 MAR 2003

USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.

PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file

10049937

provided by InfoChem.

STRUCTURE FILE UPDATES: 25 MAR 2003 HIGHEST RN 500688-79-9
DICTIONARY FILE UPDATES: 25 MAR 2003 HIGHEST RN 500688-79-9

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when
conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP
PROPERTIES for more information. See STNote 27, Searching Properties
in the CAS Registry File, for complete details:
<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

=>

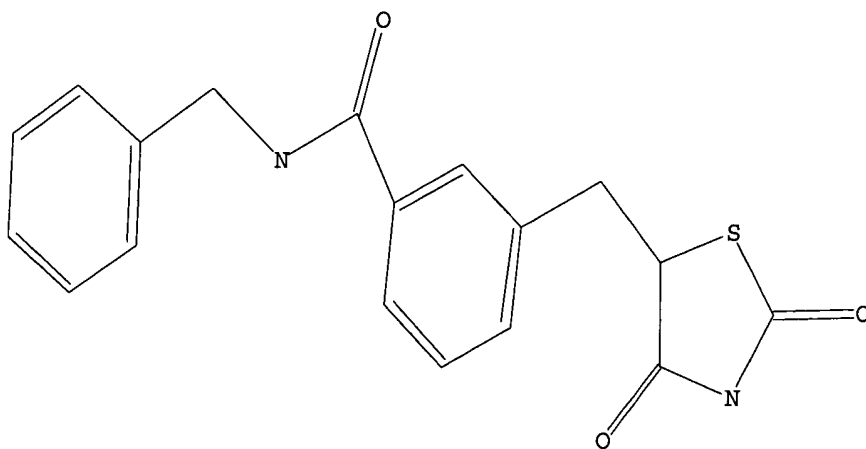
Uploading 10049937.str

L1 STRUCTURE UPLOADED

=> D

L1 HAS NO ANSWERS

L1 STR



Structure attributes must be viewed using STN Express query preparation.

=> S L1 SSS SAM

SAMPLE SEARCH INITIATED 08:57:50 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED - 13 TO ITERATE

100.0% PROCESSED 13 ITERATIONS

2 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**

PROJECTED ITERATIONS: 44 TO 476

PROJECTED ANSWERS: 2 TO 124

L2 2 SEA SSS SAM L1

10049937

=> S L1 FULL

FULL SEARCH INITIATED 08:57:54 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 213 TO ITERATE

100.0% PROCESSED 213 ITERATIONS
SEARCH TIME: 00.00.01

53 ANSWERS

L3 53 SEA SSS FUL L1

=> FIL CAPLUS

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

148.15

148.36

FILE 'CAPLUS' ENTERED AT 08:57:59 ON 26 MAR 2003
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 26 Mar 2003 VOL 138 ISS 13
FILE LAST UPDATED: 25 Mar 2003 (20030325/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> S L3 FULL

L4 42 L3

=> D L4 1-42 IBIB ABS HITSTR

L4 ANSWER 1 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:927185 CAPLUS

DOCUMENT NUMBER: 138:24716

TITLE: Preparation of azolecarboxylic acids useful as antidiabetic and antiobesity agents

INVENTOR(S): Cheng, Peter T.; Zhang, Hao; Hariharan, Narayanan

PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA

SOURCE: PCT Int. Appl., 169 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
-----	----	-----	-----	-----

WO 2002096358 A2 20021205 WO 2002-US16633 20020523

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

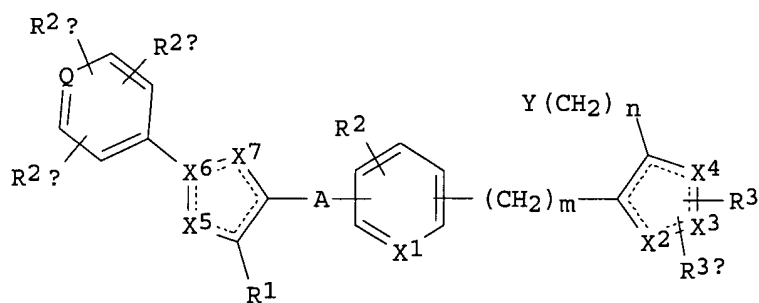
PRIORITY APPLN. INFO.:

US 2001-294380P P 20010530

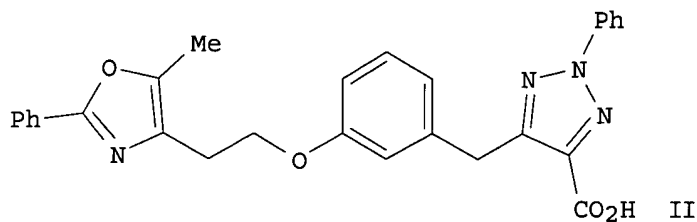
OTHER SOURCE(S):

MARPAT 138:24716

GI



I



II

AB Title compds. [I; m, n = 0-2; Q = C, N; A = (CH₂)_x, (CH₂)_{x1}, (CH₂)_{x20}(CH₂)_{x3}; x = 1-5; x₁ = 2-5; x₂, x₃ = 0-5; .gtoreq.1 of x₂, x₃ .noteq. 0; X₁ = CH, N; X₂, X₃, X₄, X₅, X₇ = C, N, O, S; in each of X₁-X₇, C may include CH; R₁ = H, alkyl; R₂ = H, alkyl, alkoxy, halo, (substituted) amino; R_{2a}, R_{2b} and R_{2c} = H, alkyl, alkoxy, halo, (substituted) amino; R₃, R_{3a} = H, alkyl, arylalkyl, aryloxy, carbonyl, alkyloxy, carbonyl, alkynyloxy, carbonyl, alkenyloxy, carbonyl, aryl, carbonyl, alkyl, carbonyl, aryl, heteroaryl, alkyl(halo)aryloxy, carbonyl, alkoxy(halo)aryloxy, carbonyl, cycloalkylaryloxy, carbonyl, cycloalkyloxyaryloxy, carbonyl, cycloheteroalkyl, heteroaryl, carbonyl, heteroaryl, heteroarylalkyl, alkyl, carbonyl, amino, aryl, carbonyl, amino, heteroaryl, carbonyl, amino, alkoxy, carbonyl, amino, aryloxy, carbonyl, amino, heteroaryl, heteroaryl, carbonyl, alkyl, sulfonyl, alkenyl, sulfonyl, heteroaryl, aryloxy, carbonyl, cycloheteroalkyloxy, carbonyl, heteroaryl, alkyl, aminocarbonyl, substituted aminocarbonyl, alkyl, aminocarbonyl, aryl, aminocarbonyl, aryloxy, arylalkyl, alkynyloxy, carbonyl, haloalkoxyaryloxy, carbonyl, alkoxy, carbonyl, aryloxy, carbonyl, aryloxy, aryloxy, carbonyl, aryl, sulfinyl, aryl, carbonyl, etc.; Y = CO₂R₄,

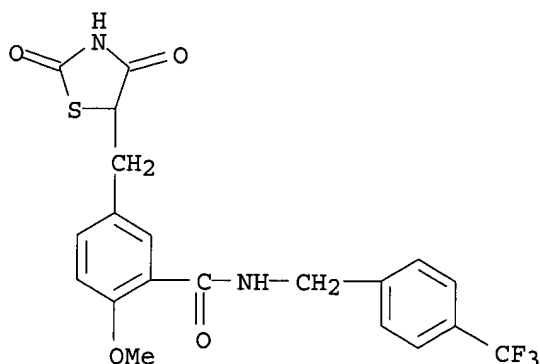
1-tetrazolyl, P(O)(OR4a)R5, P(O)(OR4a)2; R4 = H, alkyl, prodrug ester; R4a = H, prodrug ester; R5 = alkyl, aryl; with provisos], were prepd. as simultaneous inhibitors of peroxisome proliferator activated receptor-.gamma. (PPAR.gamma.) and stimulators of peroxisome proliferator activated receptor-.alpha. (PPAR.alpha.). Thus, title compd. (II) (prepd. starting from Meldrum's acid 3-methoxyphenylacetyl chloride) bound to human PPAR.alpha. and to PPAR.gamma. ligand binding domains with IC50 = 69 nM.

IT 213252-19-8, Krp297

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (coadministration; prepn. of azolecarboxylic acids useful as antidiabetic and antiobesity agents)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 2 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:927184 CAPLUS

DOCUMENT NUMBER: 138:14048

TITLE: Preparation of oxazolyloxyphenylprolines and related compounds as antidiabetic and antiobesity agents.

INVENTOR(S): Cheng, Peter T.; Jeon, Yoon; Wang, Wei

PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA

SOURCE: PCT Int. Appl., 107 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002096357	A2	20021205	WO 2002-US16628	20020523
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,			

CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR,
BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

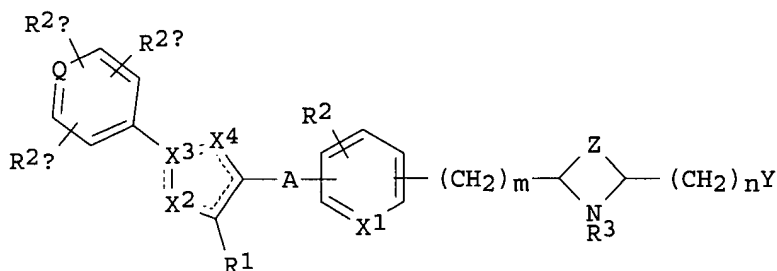
PRIORITY APPLN. INFO.:

US 2001-294505P P 20010530

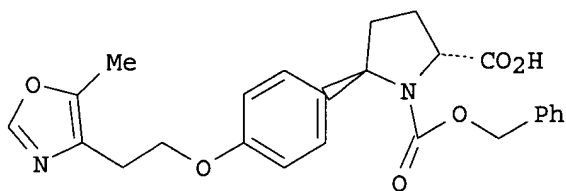
OTHER SOURCE(S):

MARPAT 138:14048

GI



I



II

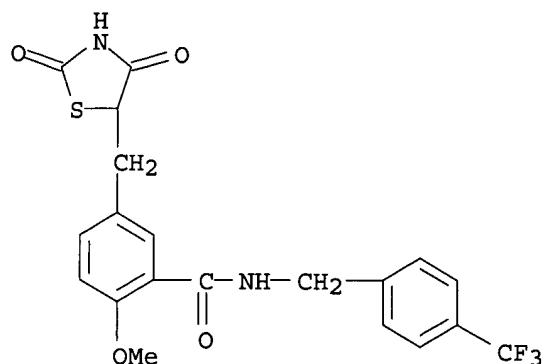
AB Title compds. [I; m, n = 0-2; Q = C, N; A = (CH₂)_x, (CH₂)_{x1}, with an alkenyl or alkynyl bond in the chain, (CH₂)_{x2}(CH₂)_{x3}; x = 1-5; x1 = 2-5; x2, x3 = 0-5; provided that .gtoreq.1 of x2 and x3 .noteq. 0; X1 = CH, N; X2 = C, N, O, S; X3 = C, N; X4 = C, N, O, S provided that .gtoreq.1 of X2, X3, X4 = N; in each of X1-X4, C may include CH; R1 = H, alkyl; R2 = H, alkyl, alkoxy, halo, (substituted) amino; R2a, R2b R2c = H, alkyl, alkoxy, halo, (substituted) amino; R3 = H, alkyl, arylalkyl, aryloxycarbonyl, alkyloxycarbonyl, alkynyloxycarbonyl, alkenyloxycarbonyl, arylcarbonyl, alkylcarbonyl, aryl, heteroaryl, cycloheteroalkyl, heteroarylcarbonyl, heteroarylheteroarylalkyl, alkylcarbonylamino, arylcarbonylamino, heteroarylcarbonylamino, alkoxy carbonylamino, aryloxycarbonylamino, heteroaryloxycarbonylamino, heteroarylheteroarylcarbonyl, alkylsulfonyl, alkenylsulfonyl, heteroaryloxycarbonyl, cycloheteroalkyloxycarbonyl, aryloxyheteroarylalkyl, heteroarylalkyloxyarylalkyl, arylarylalkyl, arylalkenylarylalkyl, arylaminoarylalkyl, etc.; Y = CO₂R₄, 1-tetrazolyl, P(O)(OR_{4a})R₅, P(O)(OR_{4a})₂; R₄ = H, alkyl, prodrug ester; R_{4a} = H, prodrug ester; R₅ = alkyl, aryl; Z = (CH₂)_{x4}, (CH₂)_{x5}, (CH₂)_{x6}(CH₂)_{x7}; x₄ = 1-5; x₅ = 2-5; x₆, x₇ = 0-4], were prepd. as antidiabetic and antiobesity agents (no data). Thus, title compd. (II) was prepd. in 6 steps.

IT 213252-19-8, KRP297

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(coadministration; prepn. of oxazolyloxyphenylprolines and related compds. as antidiabetic and antiobesity agents)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 3 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:889852 CAPLUS

DOCUMENT NUMBER: 137:346174

TITLE: Use of PPAR-.gamma.agonists for the prevention or treatment of diseases associated with IL-10 production

INVENTOR(S): Winiski, Anthony

PATENT ASSIGNEE(S): Novartis AG, Switz.

SOURCE: Brit. UK Pat. Appl., 14 pp.

CODEN: BAXXDU

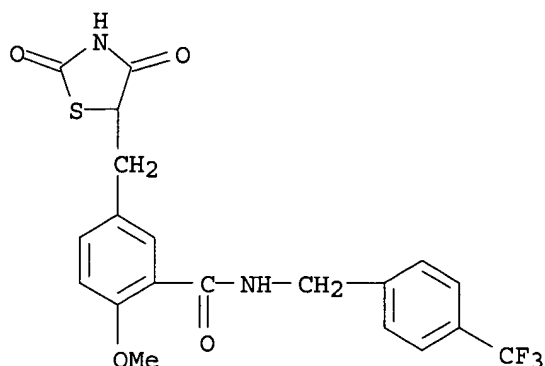
DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

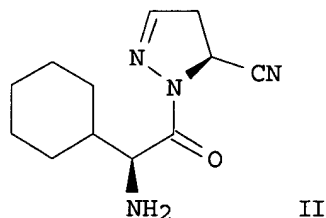
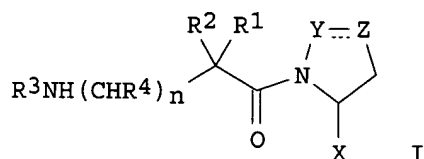
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
GB 2373725	A1	20021002	GB 2001-8087	20010330
PRIORITY APPLN. INFO.:			GB 2001-8087	20010330
AB The invention discloses the use of PPAR-.gamma. agonists for the treatment of diseases related to the prodn. of Interleukin-10 (IL-10) like systemic lupus erythematosus, arthritis, cancer etc.				
IT 213252-19-8, KRP-297				
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)				
(use of PPAR gamma agonist for prevention or treatment of diseases assocd. with IL-10 prodn.)				
RN 213252-19-8 CAPLUS				
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)				



L4 ANSWER 4 OF 42 CAPLUS COPYRIGHT 2003 ACS
 ACCESSION NUMBER: 2002:813924 CAPLUS
 DOCUMENT NUMBER: 137:311200
 TITLE: Preparation of 2,1-oxazoline and 1,2-pyrazoline-based inhibitors of dipeptidyl peptidase IV
 INVENTOR(S): Sulsky, Richard B.; Robl, Jeffrey A.
 PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA
 SOURCE: PCT Int. Appl., 61 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002083128	A1	20021024	WO 2002-US10936	20020405
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
US 2002183367	A1	20021205	US 2002-107279	20020326
PRIORITY APPLN. INFO.: US 2001-283438P P 20010412 OTHER SOURCE(S): MARPAT 137:311200 GI				



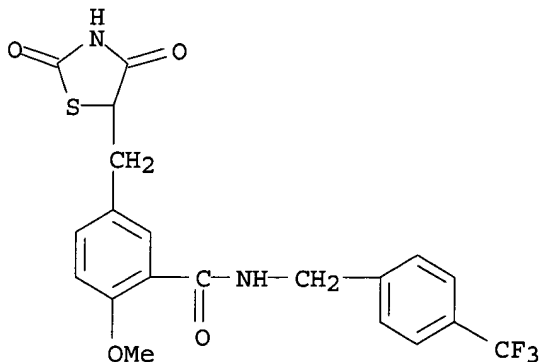
AB The invention describes dipeptidyl peptidase IV (DP 4) inhibiting compds. I [n is 0 or 1; X is H or CN; Y is N, NH or O; Z is CH₂ when Y is O or NH, with Y-Z forming a single bond, and Z is CH when Y is N, with Y-Z forming a double bond; R1-R4 = H, alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkylalkyl, bicycloalkyl, bicycloalkylalkyl, alkylthioalkyl, arylalkylthioalkyl, cycloalkenyl, aryl, aralkyl, heteroaryl, heteroarylalkyl, cycloheteroalkyl or cycloheteroalkylalkyl, which may be substituted; R1 may combine with R3 or R4 to form a ring (CR₅R₆)₂₋₆ or (CR₇R₈)₃₋₆, resp., where R₅-R₈ = H, OH, alkoxy, alkyl, aryl, etc.] and their pharmaceutically-acceptable salts or prodrug esters. A method is also provided for treating diabetes and related diseases, employing a DP 4 inhibitor I, optionally in combination with other therapeutic agents, including an antidiabetic, hypolipidemic, or anti-obesity agent. Thus, coupling of sultam-protected 1,2-pyrazoline-3-carboxamide with (S)-N-(tert-butoxycarbonyl)cyclohexylglycine (HOAt, Et₃N, and EDAC in CH₂Cl₂), followed by sultam cleavage with methanolic ammonia, amide conversion to nitrile using imidazole, and deprotection, afforded II.TFA.

IT 213252-19-8, KRP297

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(antidiabetic agent; prepn. of oxazoline and pyrazoline-based inhibitors of dipeptidyl peptidase IV)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

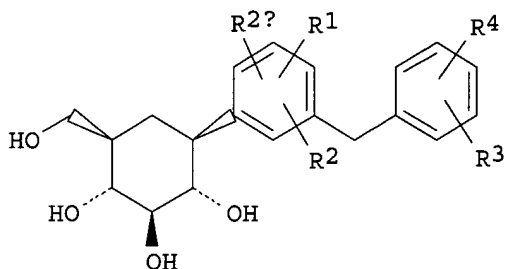
L4 ANSWER 5 OF 42 CAPLUS COPYRIGHT 2003 ACS
 ACCESSION NUMBER: 2002:813874 CAPLUS
 DOCUMENT NUMBER: 137:311199
 TITLE: Amino acid complexes of C-aryl glucosides for treatment of diabetes
 INVENTOR(S): Gougoutas, Jack Z.
 PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA
 SOURCE: PCT Int. Appl., 80 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002083066	A2	20021024	WO 2002-US11066	20020408
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				

PRIORITY APPLN. INFO.: US 2001-283097P P 20010411

OTHER SOURCE(S): MARPAT 137:311199

GI



I

AB Cryst. complexes are obtained from 1:1 or 2:1 mixts. of either the (D) or (L) enantiomer of natural amino acids and compds. of formula I [R1, R2, R2a = H, OH, OR5, alkyl, OCHF2, OCF3, SR5a, halogen; R3, R4 = H, OH, OR5b, alkyl, cycloalkyl, CF3, OCHF2, OCF3, halogen, CONR6R6a, CO2R5c, CO2H, COR6b, CH(OH)R6c, CH(OR5d)R6d, CN, NHCOR5e, NHSO2R5f, NHSO2-aryl, SR5g, SOR5h, SO2R5i, or a five, six or seven membered heterocycle which may contain 1 to 4 heteroatoms (N, O, S, SO, and/or SO2), or R3 and R4 together with the carbons to which they are attached form an annelated five, six or seven membered carbocycle or heterocycle which may contain 1 to 4 heteroatoms in the ring; R5, R5a-R5i are independently alkyl; R6,

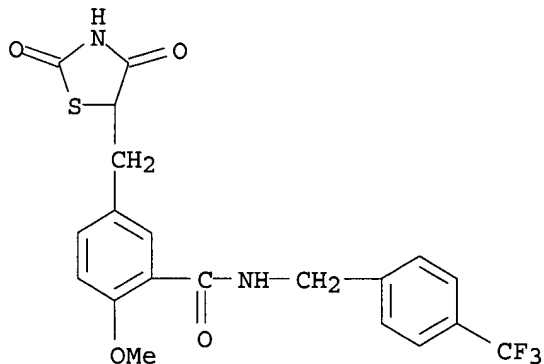
R6a-R6d are independently H, alkyl, aryl, alkylaryl or cycloalkyl, or NR6R6a form an annelated five, six or seven membered heterocycle which may contain 1 to 4 heteroatoms in the ring]. A method is also provided for treating diabetes and related diseases employing an SGLT2 (sodium dependent glucose transporters found in the intestine and kidney) inhibiting amt. of the above complex alone or in combination with another antidiabetic agent or other therapeutic agent. Thus, I (R1 = 4-Me, R4 = 4-OCHF2, R2, R2a, R3 = H) was prepd. by a multistep procedure starting from o-toluic acid, anisole, 2,3,4,6-tetra-O-benzyl-.beta.-D-glucolactone, and CHF2Cl and treated with L-phenylalanine to form the cryst. 1:1 complex.

IT 213252-19-8, KRP297

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(prepn. of amino acid/C-aryl glucoside complexes for treatment of diabetes and related diseases)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 6 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:793435 CAPLUS

DOCUMENT NUMBER: 137:289021

TITLE: Combination therapy comprising glucose reabsorption inhibitors and PPAR modulators

INVENTOR(S): Bussolari, Jacqueline C.; Chen, Xiaoli; Conway, Bruce R.; Demarest, Keith T.; Ross, Hamish N. M.; Severino, Rafael

PATENT ASSIGNEE(S): Ortho McNeil Pharmaceutical, Inc., USA

SOURCE: PCT Int. Appl., 86 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002080936	A1	20021017	WO 2002-US10538	20020403
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,				

PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
 UA, UG, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,
 CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR,
 BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

US 2003045553 A1 20030306 US 2002-115827 20020403

PRIORITY APPLN. INFO.: US 2001-281429P P 20010404

AB Combination therapy comprising PPAR modulators and glucose reabsorption inhibitors useful for the treatment of diabetes and Syndrome X are disclosed.

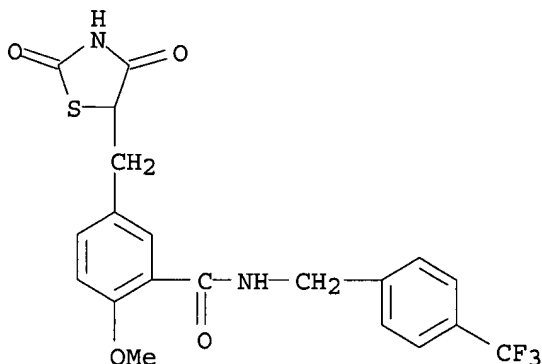
IT 213252-19-8, KRP-297

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
 (Biological study); USES (Uses)

(combination therapy comprising glucose reabsorption inhibitors and PPAR modulators)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 7 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:736927 CAPLUS

DOCUMENT NUMBER: 137:247879

TITLE: Preparation of antidiabetic agents C-aryl glucoside as human SGLT2 inhibitors

INVENTOR(S): Ellsworth, Bruce; Washburn, William N.; Sher, Philip M.; Wu, Gang; Meng, Wei

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 17 pp., Cont.-in-part of U.S. 6,414,126.

CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2002137903	A1	20020926	US 2002-151436	20020520
US 6515117	B2	20030204		
US 6414126	B1	20020702	US 2000-679027	20001004

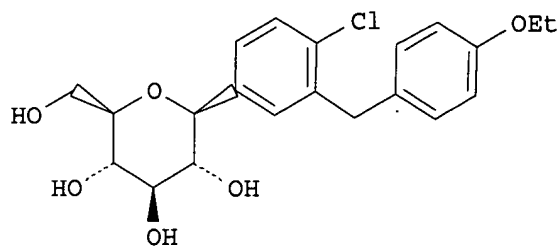
PRIORITY APPLN. INFO.:

US 1999-158773P P 19991012

US 2000-194615P P 20000405

US 2000-679027 A2 20001004

GI



I

AB An SGLT2 inhibiting compd. is provided having the formula I method is also provided for treating diabetes and related diseases employing an SGLT2 inhibiting amt. of the above compd. alone or in combination with another antidiabetic agent or other therapeutic agent (no data). 1A pharmaceutical combination comprising an SGLT2 inhibitor compd. and an antidiabetic agent other than an SGLT2 inhibitor, for treating the complications of diabetes, an anti-obesity agent, an antihypertensive agent, an antiplatelet agent, an antiatherosclerotic agent, and/or a lipid-lowering agent (no data). A method for treating or delaying the progression or onset of diabetes, diabetic retinopathy, diabetic neuropathy, diabetic nephropathy, delayed wound healing, insulin resistance, hyperglycemia, hyperinsulinemia, elevated blood levels of fatty acids or glycerol, hyperlipidemia, obesity, hypertriglyceridemia, Syndrome X, diabetic complications, atherosclerosis or hypertension, or for increasing high d. lipoprotein levels, which comprises administering to a mammalian species in need of treatment a therapeutically effective amt. of a compd (no data).

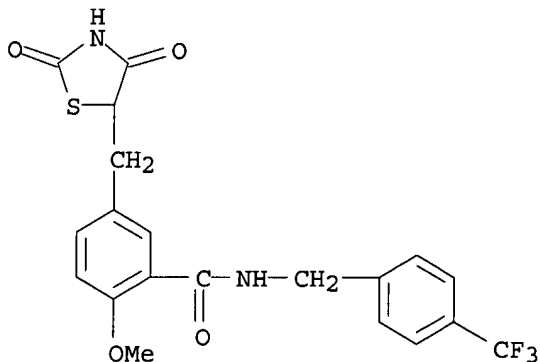
IT 213252-19-8, KRP297

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(prepn. of antidiabetic agents C-aryl glucosides as human SGLT2 inhibitors)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 8 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:637483 CAPLUS

DOCUMENT NUMBER: 137:185311

TITLE: Preparation of 2-aryloxy-2-arylalkanoic acids for diabetes and lipid disorders

INVENTOR(S): Adams, Alan D.; Jones, A. Brian; Berger, Joel P.; Dropinski, James F.; Elbrecht, Alexander; Liu, Kun; Macnaul, Karen Lamb; Shi, Guo-qiang; Von, Langen Derek J.; Zhou, Gaochao

PATENT ASSIGNEE(S): Merck & Co., Inc., USA

SOURCE: PCT Int. Appl., 157 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

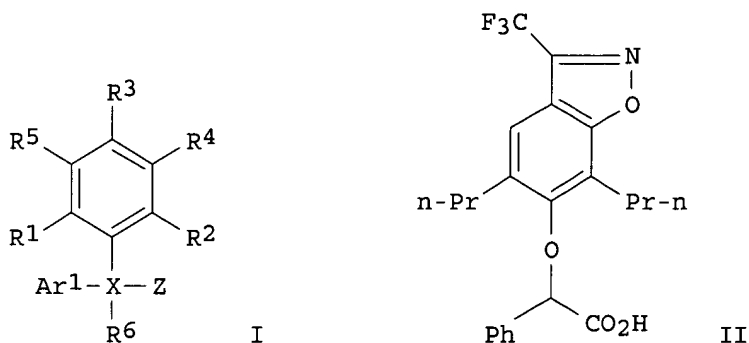
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002064094	A2	20020822	WO 2002-US4680	20020205
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				

PRIORITY APPLN. INFO.: US 2001-267809P P 20010209

OTHER SOURCE(S): MARPAT 137:185311

GI



AB Title compds. I [R1 = halo, alkyl, alkoxy; R2 = alkyl, alicyclic; R3 = alkyl, aryl, alicyclic, heterocycle, etc.; R4 = H, OH, alkoxy, aryloxy, halo or R3-4 may be joined together to yield 5- or 6-membered heterocycle; R5 = H, halo; R6 = H, halo, CH3, CF3; Ar1 = Ph, thienyl, thiazolyl, oxazolyl, pyridyl; X = O, S; Z = COOH, tetrazole, carboxamide] were prepd. For instance, 2,4-dipropylresorcinol was converted to 2,4-dihydroxy-3,5-dipropyl-.alpha.,.alpha.,.alpha.-trifluoroacetophenone (CH2Cl2, TFAA,

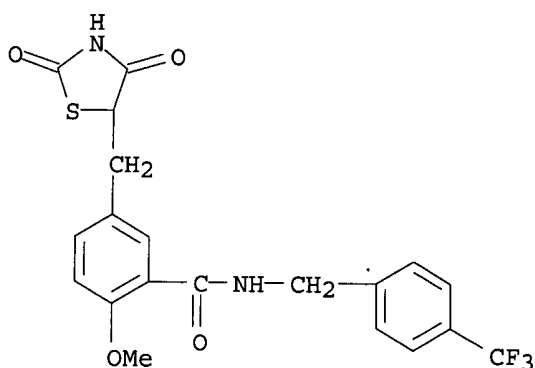
AlCl₃) and subsequently treated with i. hydroxylamine.bul.HCl, MeOH, reflux; ii. Ac₂O; iii. pyridine, reflux which afforded 5,7-dipropyl-6-hydroxy-3-trifluoromethyl-1,2-benzisoxazole. The benzisoxazole was reacted with Me 2-bromo-2-phenylacetate (DMF, Cs₂CO₃) and the product saponified to give II. I are potent agonists of the peroxisome proliferator activated receptor and are useful in the treatment of non-insulin dependent diabetes mellitus (NIDDM), hyperglycemia, dyslipidemia, hyperlipidemia, hypercholesterolemia, hypertriglyceridemia, atherosclerosis, obesity, vascular restenosis, inflammation, and other PPAR-.alpha. and/or PPAR-.gamma. mediated diseases.

IT 213252-19-8, KRP-297

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(combination pharmaceutical; prepn. of 2-aryloxy-2-arylalkanoic acids for diabetes and lipid disorders)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 9 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:594636 CAPLUS

DOCUMENT NUMBER: 137:135097

TITLE: Acyl sulfamides for treatment of obesity, diabetes and lipid disorders

INVENTOR(S): Jones, A. Brian; Acton, John J., III

PATENT ASSIGNEE(S): Merck & Co., Inc., USA

SOURCE: PCT Int. Appl., 64 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002060388	A2	20020808	WO 2002-US3119	20020125
WO 2002060388	A3	20030227		

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,
CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR,
BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.: US 2001-264955P P 20010130

OTHER SOURCE(S): MARPAT 137:135097

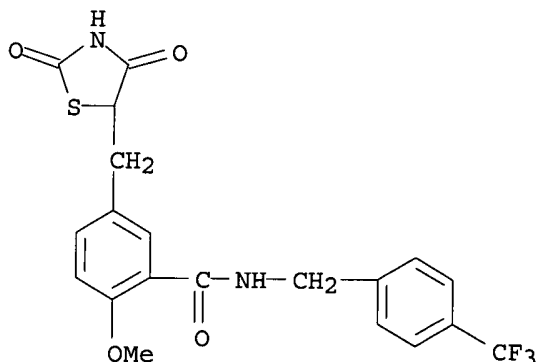
AB A class of acyl sulfamides comprises compds. that are potent ligands for PPAR.gamma. receptors and generally have antagonist or partial agonist activity. The compds. may be useful in the treatment, control or prevention of obesity, non-insulin dependent diabetes mellitus (NIDDM), hyperglycemia, dyslipidemia, hyperlipidemia, hypercholesterolemia, hypertriglyceridemia, atherosclerosis, vascular restenosis, inflammation, and other PPAR.gamma. receptor-mediated diseases, disorders and conditions, alone or in combination with one or more other compds. Other compds. are selected from insulin sensitizers, insulin or insulin mimetics, sulfonylureas, .alpha.-glucosidase inhibitors, cholesterol lowering agents, PPAR.delta. agonists, antiobesity compds., an ileal bile acid transporter inhibitor, and agents intended for use in inflammatory conditions such as aspirin, nonsteroidal anti-inflammatory drugs, glucocorticoids, azulfidine, and cyclooxygenase-2 selective inhibitors.

IT 213252-19-8, KRP-297

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(acyl sulfamides and other drugs for treatment of metabolic disorders mediated by PPAR.gamma. receptors)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 10 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:575765 CAPLUS

DOCUMENT NUMBER: 137:140435

TITLE: Benzopyrancarboxylic acid derivatives with PPAR agonist activity for the treatment of diabetes and lipid disorders, and their preparation, pharmaceutical compositions, and use

INVENTOR(S): Sahoo, Soumya P.; Koyama, Hiroo; Miller, Daniel J.; Boueres, Julia K.; Desai, Ranjit C.

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 42 pp.

CODEN: USXXCO

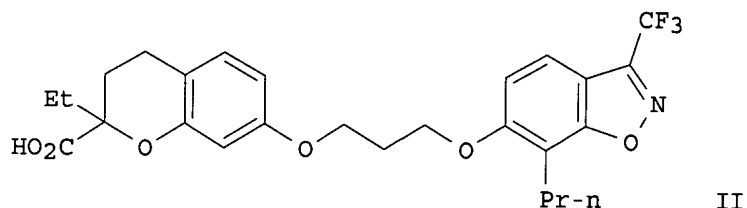
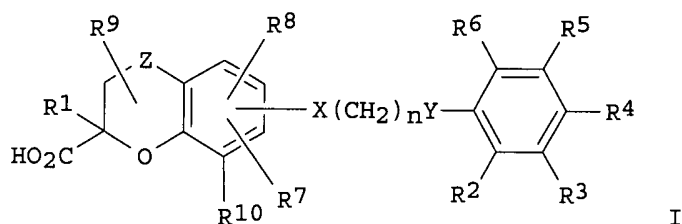
DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2002103242	A1	20020801	US 2001-21667	20011029
WO 2002060434	A2	20020808	WO 2001-US49501	20011026
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
PRIORITY APPLN. INFO.:			US 2000-244698P P 20001031	
OTHER SOURCE(S):		MARPAT 137:140435		
GI				



AB A class of benzopyrancarboxylic acid derivs. is disclosed, which comprises compds. that are potent agonists (no data) of peroxisome proliferator activated receptors (PPAR) alpha and/or gamma, and are therefore useful in the treatment, control, or prevention of non-insulin dependent diabetes mellitus (NIDDM), hyperglycemia, dyslipidemia, hyperlipidemia, hypercholesterolemia, hypertriglyceridemia, atherosclerosis, obesity, vascular restenosis, inflammation, and other PPAR alpha and/or gamma mediated diseases, disorders and conditions. In particular, compds. I and their pharmaceutically acceptable salts and/or prodrugs are disclosed [wherein: Z = CH₂, CO; R₁ = H, OH, halo, (un)substituted alk(en/yn)yl, alk(en/yn)yloxy, or aryl; or R₁ forms (un)substituted cyclopropane fusion to adjacent C atom; X, Y = O, S, SO, SO₂, CH₂, (un)substituted NH; n = 1-6; R₄ = (un)substituted benzoheterocyclyl, cycloalkyl, heterocyclyl, cycloalkyloxy, halo, OH or derivs., alk(en/yn)yl, alk(en/yn)yloxy, or aryl, etc.; other R groups = H, halo, OH, (un)substituted alk(en/yn)yl, alk(en/yn)yloxy, aryl, aryloxy, aroyl, etc.; or R₃R₄ or R₄R₅ = (un)substituted 5- or 6-membered heterocyclic ring]. A list of 29 compds.

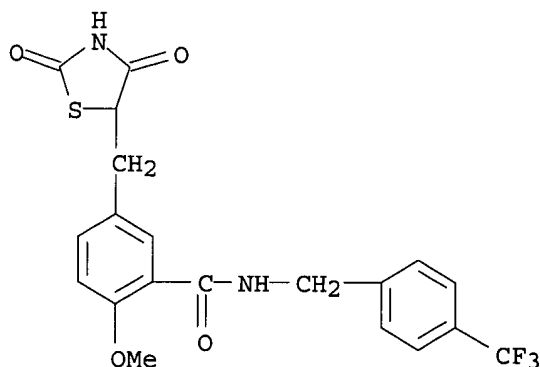
is claimed, and their prepn. is described. For example, Et 7-hydroxy-4-oxo-4H-chromene-2-carboxylate underwent a sequence of: (1) complete hydrogenation of the enone (98%), (2) etherification of the alc. with PhCH₂O(CH₂)₃Br (66%), (3) alpha ethylation of the ester (70%), (4) hydrogenolytic debenzoylation (100%), (5) conversion of the resultant alc. to a bromide (96%), (6) etherification of the bromide with 3-(trifluoromethyl)-7-propyl-6-hydroxybenz[4,5]isoxazole (85%), and (7) alk. hydrolysis (100%), to give title compd. II. PPAR binding assays using human recombinant PPAR are described without data. Co-administration of compds. I with a variety of other drug categories, including a no. of specific drugs, is claimed.

IT 213252-19-8, KRP-297

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(therapeutic compns. also contg.; prepn. of benzopyrancarboxylic acid derivs. as PPAR agonists for treatment of diabetes and lipid disorders)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 11 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:540258 CAPLUS

DOCUMENT NUMBER: 137:109267

TITLE: Preparation of benzoxepinopyridines as HMG-CoA reductase inhibitors

INVENTOR(S): Robl, Jeffrey A.; Chen, Bang-chi; Sun, Chong-qing

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 42 pp., Cont.-in-part of U.S. Ser. No. 875,155.

CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2002094977	A1	20020718	US 2001-7407	20011204
US 2002013334	A1	20020131	US 2001-875155	20010606
PRIORITY APPLN. INFO.:			US 2000-211595P	P 20000615
			US 2001-875155	A2 20010606

OTHER SOURCE(S): MARPAT 137:109267

GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

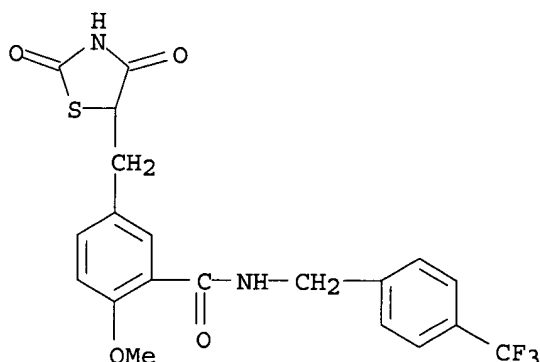
AB Title compds. I [X = O, S, SO, SO₂, NR₇; Z = HOCHCH₂CH(OH)CH₂CO₂R₃, 4-hydroxy-2-oxopyran-6-yl, etc.; n = 0, 1; R₁, R₂ = alkyl, arylalkyl, cycloalkyl, alkenyl, cycloalkenyl, aryl, heteroaryl, cycloheteroalkyl; R₃ = H, alkyl, metal ion; R₄ = H, halo, CF₃, etc.; R₇ = H, alkyl, aryl, alkanoyl, aroyl, alkoxycarbonyl, etc.; R₉, R₁₀ = H, alkyl], were prepd. as HMG CoA reductase inhibitors active in inhibiting cholesterol biosynthesis, modulating blood serum lipids such as lowering LDL cholesterol and/or increasing HDL cholesterol, and treating hyperlipidemia, hypercholesterolemia, hypertriglyceridemia and atherosclerosis (no data). A multistep synthesis of II is reported.

IT 213252-19-8, KRP297

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(coadministered agents; prepn. of benzoxepinopyridines as HMG-CoA reductase inhibitors for treatment of hyperlipidemia, hypercholesterolemia, hypertriglyceridemia, atherosclerosis, and other disorders)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 12 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:504648 CAPLUS

DOCUMENT NUMBER: 137:83637

TITLE: Medicinal compositions containing diuretic and insulin resistance-improving agent

INVENTOR(S): Takaoka, Masaya; Araki, Kazushi; Kanda, Shoichi

PATENT ASSIGNEE(S): Sankyo Company, Limited, Japan

SOURCE: PCT Int. Appl., 183 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

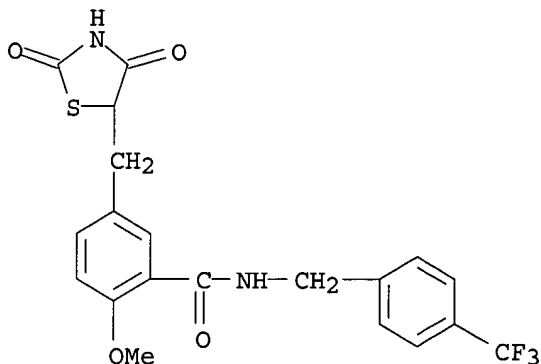
PATENT INFORMATION:

PATENT NO.

KIND DATE

APPLICATION NO. DATE

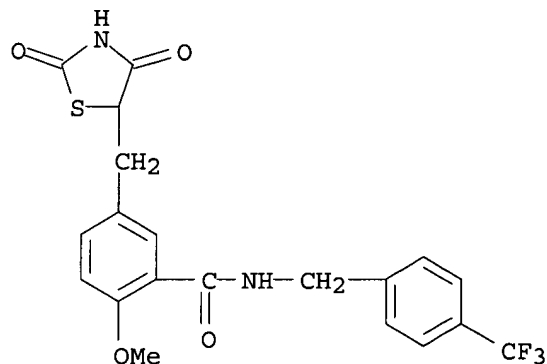
 WO 2002051441 A1 20020704 WO 2001-JP11296 20011221
 W: AU, BR, CA, CN, CO, CZ, HU, ID, IL, IN, KR, MX, NO, NZ, PH, PL,
 RU, SG, SK, US, VN, ZA
 RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
 PT, SE, TR
 JP 2002255854 A2 20020911 JP 2001-386861 20011220
 PRIORITY APPLN. INFO.: JP 2000-394424 A 20001226
 OTHER SOURCE(S): MARPAT 137:83637
 AB Disclosed are medicinal compns. contg. a diuretic and an insulin
 resistance-improving agent whereby side effects assocg. the administration
 of an insulin resistance-improving agent (for example, megalocardia,
 edema, body fluid retention, pleural effusion) can be prevented or
 treated. Oral administration of furosemide prevented increases of heart
 wt. and blood plasma, and edema due to administration of
 5-[4-(6-methoxy-1-methyl-1H-benzimidazol-2-ylmethoxy)benzyl]thiazolidine-
 2,4-dione hydrochloride.
 IT 213252-19-8, KRP-297
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (medicinal compns. contg. diuretics and insulin resistance-improving
 agents)
 RN 213252-19-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-
 (trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 22 THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS
 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 13 OF 42 CAPLUS COPYRIGHT 2003 ACS
 ACCESSION NUMBER: 2002:409256 CAPLUS
 DOCUMENT NUMBER: 137:735
 TITLE: Methods and compositions for treatment of diabetes and
 related conditions via gene therapy
 INVENTOR(S): Caplan, Shari L.; Boettcher, Brian R.; Slosberg, Eric
 D.; Connelly, Sheila; Kaleko, Michael; Desai, Urvi J.
 PATENT ASSIGNEE(S): USA
 SOURCE: U.S. Pat. Appl. Publ., 42 pp.
 CODEN: USXXCO
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	-----	----	-----	-----	-----
	US 2002065239	A1	20020530	US 2001-808457	20010314
PRIORITY APPLN. INFO.:				US 2000-266328P P	20000315
AB	Methods and compns. are disclosed for the treatment of diabetes, obesity and diabetic-related conditions. The methods include gene therapy based administration of a therapeutically effective amt. of vectors encoding the following: glucokinase regulatory protein alone or co-administered with glucokinase or with metab. modifying proteins; glucokinase co-administered with metab. modifying proteins; or glucokinase regulatory protein co-administered with glucokinase in combination with metab. modifying proteins, to a diabetic patient. The metab. modifying proteins include UCP2, UCP3, PPAR.alpha., OB-Rb, GLP-1 and GLP-1 analogs (administered via vector or directly as a peptide). Preferred examples of GLP-1 analogs include GLP-1-Gly8, Exendin-4 and the "Black Widow" chimeric GLP-1 analog. Addnl., PPAR.alpha. ligands and DPP-IV inhibitors may be co-administered with the above.				
IT	213252-19-8, KRP-297				
	RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)				
	(gene therapy for treatment of diabetes and related conditions)				
RN	213252-19-8 CAPLUS				
CN	Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)				



L4 ANSWER 14 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:392237 CAPLUS

DOCUMENT NUMBER: 136:401651

TITLE: Preparation of fused pyridine derivatives as HMG-CoA reductase inhibitors

INVENTOR(S): Robl, Jeffrey A.; Chen, Bang-Chi; Sun, Chong-Qing

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 46 pp., Cont.-in-part of U.S. Ser. No. 875,218.

CODEN: USXXCO

DOCUMENT TYPE: Patent

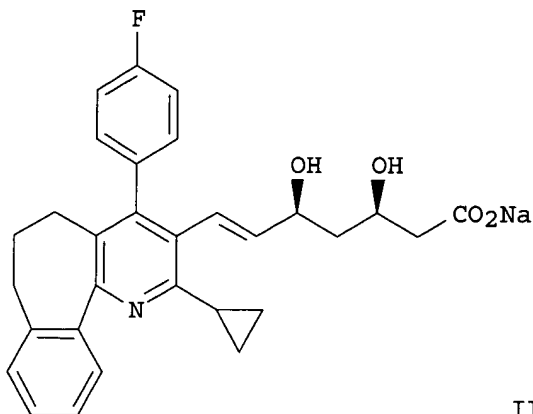
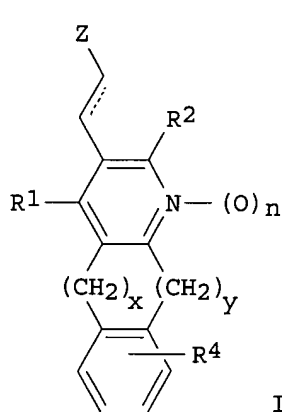
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
-----	----	-----	-----	-----

US 2002061901 A1 20020523 US 2001-8154 20011204
 US 2002028826 A1 20020307 US 2001-875218 20010606
 PRIORITY APPLN. INFO.: US 2000-211594P P 20000615
 US 2001-875218 A2 20010606
 OTHER SOURCE(S): MARPAT 136:401651
 GI



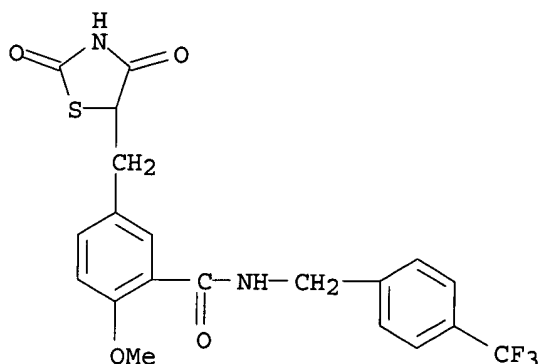
AB The title compds. I and their pharmaceutically acceptable salts, esters, prodrug esters, and stereoisomers are claimed [wherein: Z = CH(OH)CH2CR7(OH)CH2CO2R3 or corresponding pyranone lactone derivs.; n = 0, 1; x = 0, 1, 2, 3, or 4; y = 0, 1, 2, 3 or 4, provided that at least one of x and y is other than 0; and optionally one or more carbons of (CH2)x and/or (CH2)y together with addnl. carbons form a 3 to 7 membered spirocyclic ring; R1, R2 = alkyl, arylalkyl, cycloalkyl, alkenyl, cycloalkenyl, aryl, heteroaryl, cycloheteroalkyl; R3 = H or lower alkyl; R4 = H, halo, CF3, OH, alkyl, alkoxy, CO2H, (un)substituted NH2, cyano, (un)substituted CONH2, etc.; R7 = H, alkyl]. The compds. are HMG-CoA reductase inhibitors, and are active in inhibiting cholesterol biosynthesis and modulating blood serum lipids, for example, lowering LDL cholesterol and/or increasing HDL cholesterol (no data). I are thus useful in treating hyperlipidemia and dyslipidemia, in hormone replacement therapy, and in treating hypercholesterolemia, hypertriglyceridemia and atherosclerosis, as well as Alzheimer's disease and osteoporosis. Prepn. of several compds. are described. For instance, a multistep synthesis of fused pyridine deriv. II is reported. Compds. I may be used in a manner similar to atorvastatin, pravastatin, simvastatin, etc. Combinations of compds. I with various other drugs are claimed, the latter being specified as certain pharmacol. classes, as inhibitors of specific enzymes, as (ant)agonists of specific receptors, and as numerous named drugs.

IT 213252-19-8, KRP297

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (therapeutic compns. also contg.; prepn. of fused pyridine derivs. as
 HMG-CoA reductase inhibitors)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 15 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:142553 CAPLUS

DOCUMENT NUMBER: 136:177969

TITLE: Medicinal compositions containing PPAR.gamma. agonists and RXR agonists for preventing and treating cancer

INVENTOR(S): Kurakata, Shinichi; Fujiwara, Kosaku; Shimazaki, Naomi; Fujita, Takashi

PATENT ASSIGNEE(S): Sankyo Company, Limited, Japan

SOURCE: PCT Int. Appl., 40 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002013864	A1	20020221	WO 2001-JP7037	20010815
W: AU, BR, CA, CN, CO, CZ, HU, ID, IL, IN, KR, MX, NO, NZ, PL, RU, SG, SK, US, ZA				
RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR				
JP 2002128700	A2	20020509	JP 2001-241740	20010809
AU 2001078738	A5	20020225	AU 2001-78738	20010815
PRIORITY APPLN. INFO.: JP 2000-246910 A 20000816				
WO 2001-JP7037 W 20010815				

OTHER SOURCE(S): MARPAT 136:177969

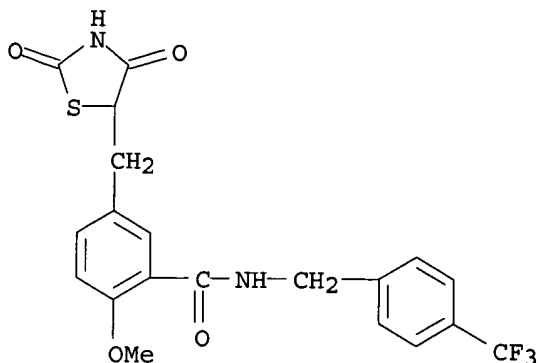
AB Disclosed are medicinal compns. for preventing or treating cancer wherein one or more Peroxisome proliferator-activated receptor .gamma. (PPAR.gamma.) activation agonists and one or more retinoid X receptor (RXR) activation agonists are used simultaneously or successively. A combined administration of 5-[4-(6-methoxy-1-methylbenzimidazol-2-ylmethoxy)benzyl]thiazolidine-2,4-dione hydrochloride (I) 5 and targretin 100 mg/kg to HL-60 cell-bearing mice showed synergistic antitumor effect. Also, tablets were prepd. from I 0.004, targretin 0.1, lactose 0.244, corn starch 50, and magnesium stearate 0.002 g.

IT 213252-19-8

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(simultaneous or successive use of PPAR.gamma. agonists and RXR agonists for prevention or treatment of cancer)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 16 OF 42 CAPLUS COPYRIGHT 2003 ACS
 ACCESSION NUMBER: 2002:142506 CAPLUS
 DOCUMENT NUMBER: 136:177977
 TITLE: Methods for treating inflammatory diseases using PPAR agonists
 INVENTOR(S): Pershadsingh, Harrihar A.
 PATENT ASSIGNEE(S): USA
 SOURCE: PCT Int. Appl., 42 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002013812	A1	20020221	WO 2001-US25668	20010816
W: AU, CA, MX, NZ, US				
RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR				
AU 2001088271	A5	20020225	AU 2001-88271	20010816
PRIORITY APPLN. INFO.:				
			US 2000-225907P	P 20000817
			US 2000-230509P	P 20000906
			WO 2001-US25668	W 20010816

AB The present invention describes methods for the use of PPAR ligands in the treatment inflammatory endocrine, dermatol., cardiovascular immunol., neurol., ophthalmic, neoplastic, pulmonary diseases, and age-related dysregulations. In addn., methods are provided for treating said conditions and diseases comprising the step of administering to a human or an animal in need thereof a therapeutic amt. of pharmacol. compns. comprising a pharmaceutically acceptable carrier, and a PPAR.gamma. agonist which cross-activates PPAR.alpha. or PPAR.delta. or both, or a PPAR.gamma. partial agonist, or a PPAR.gamma./RXR agonist, effective to reverse, slow, stop, or prevent the pathol. inflammatory or degenerative process.

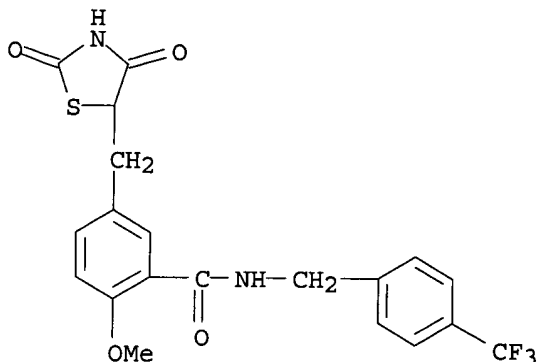
IT 213252-19-8, KRP 297

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (methods for treating inflammatory diseases using PPAR agonists)

10049937

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 17 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:90008 CAPLUS

DOCUMENT NUMBER: 136:151071

TITLE: Preparation of N-substituted indoles for treating diabetes

INVENTOR(S): Acton, John J., III; Black, Regina Marie; Jones, Anthony Brian; Wood, Harold Blair

PATENT ASSIGNEE(S): Merck & Co., Inc., USA

SOURCE: PCT Int. Appl., 73 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

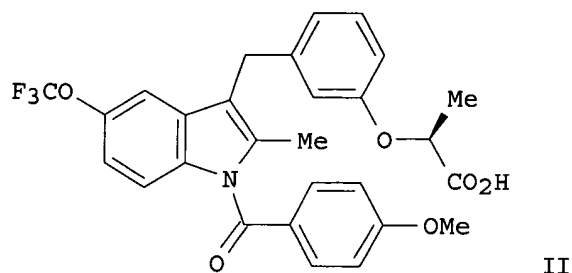
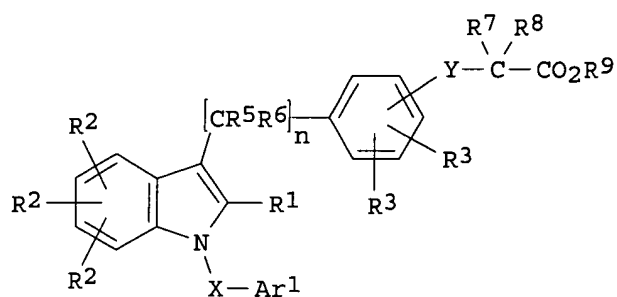
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002008188	A1	20020131	WO 2001-US22979	20010720
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
US 2002042441	A1	20020411	US 2001-912961	20010725
US 6525083	B2	20030225		

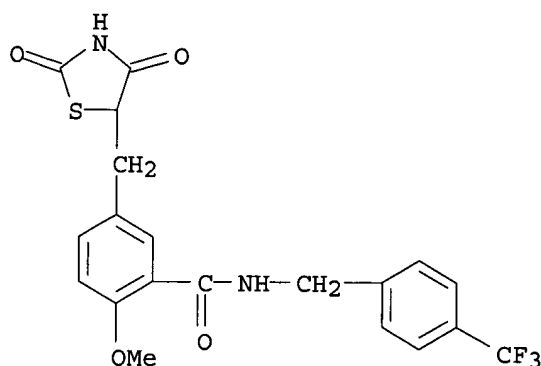
PRIORITY APPLN. INFO.: US 2000-220778P P 20000725

OTHER SOURCE(S): MARPAT 136:151071

GI



- AB The title indoles having aryloxyacetic acid substituents [I; R1 = Me, optionally substituted with 1-3 F atoms; R2-R4 = H, halo, alkyl, etc.; R5, R6 = H, F, OH, alkyl; and R5 and R6 groups that are on the same carbon atom optionally may be joined to form a cyclopropyl group; R7, R8 = H, F, alkyl; or CR7R8 may form cycloalkyl; R9 = H, alkyl; Ar1 = (un)substituted Ph, naphthyl, pyridyl, quinolyl; X = CO, SO2, CH2, CHMe, CMe2, CF2, cyclopropylidene; Y = O, S; n = 0-5] which are agonists or partial agonists of PPAR gamma, and are useful in the treatment, control or prevention of non-insulin dependent diabetes mellitus (NIDDM), hyperglycemia, dyslipidemia, hyperlipidemia, hypercholesterolemia, hypertriglyceridemia, atherosclerosis, obesity, vascular restenosis, inflammation, and other PPAR mediated diseases, disorders and conditions, were prepd. E.g., a multi-step synthesis of (2S)-II was given.
- IT 213252-19-8, KRP-297
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (prepn. of N-substituted indoles for treating diabetes)
- RN 213252-19-8 CAPLUS
- CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 18 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:56491 CAPLUS

DOCUMENT NUMBER: 137:73203

TITLE: Pharmacological analysis of wild-type .alpha., .gamma. and .delta. subtypes of the human peroxisome proliferator-activated receptor

AUTHOR(S): Wurch, T.; Junquero, D.; Delhon, A.; Pauwels, P. J.

CORPORATE SOURCE: Department of Cellular and Molecular Biology, Centre de Recherche Pierre Fabre, Castres, 81106, Fr.

SOURCE: Naunyn-Schmiedeberg's Archives of Pharmacology (2002), 365(2), 133-140

CODEN: NSAPCC; ISSN: 0028-1298

PUBLISHER: Springer-Verlag

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Three distinct peroxisome proliferator-activated receptor (PPAR) cDNAs were isolated from human brain RNA. Whereas the PPAR.delta. subtype perfectly matched the amino acid sequences reported in the Genbank database, several differences were found for the PPAR.alpha. (Lys123Met, Ala268Val, Gly296Ala and Val444Ala) and PPAR.gamma.2 (Met8Ile, Pro9Ala, Met186Ile, Pro187Ala and the deletion of a Gln213 residue) subtypes. A pharmacol. anal. was undertaken by co-expressing each PPAR subtype with a reporter plasmid contg. a luciferase gene under the transcriptional control of a synthetic, triplicated PPAR response element in either HepG2 or Cos-7 cells. Whereas fenofibrate unselectively activated the PPAR.alpha. and PPAR.delta. subtypes, the related BM-17.0744 compd. was more potent and selective for PPAR.alpha.. The thiazolidine dione derivs. rosiglitazone and pioglitazone were potent and selective PPAR.gamma.2 agonists. L-165041, reported as a selective and potent PPAR.delta. ligand, displayed in this specified transactivation system, apart from its highly efficacious PPAR.delta. agonist activity, partial and full agonism at, resp., PPAR.alpha. and PPAR.gamma.2 subtypes. In conclusion, transcriptional control of a luciferase gene by wild-type PPAR subtypes provides powerful recombinant assays to evaluate ligand's efficacy at these nuclear receptors.

IT 213252-19-8, KRP-297

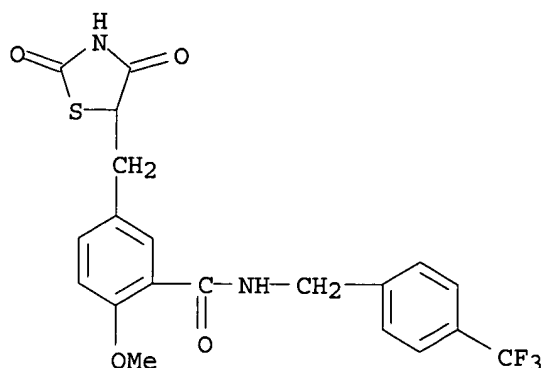
RL: PAC (Pharmacological activity); BIOL (Biological study)

(pharmacol. anal. of wild-type .alpha., .gamma. and .delta. subtypes of human peroxisome proliferator-activated receptor)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-

(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 40 THERE ARE 40 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 19 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2001:900080 CAPLUS

DOCUMENT NUMBER: 136:318816

TITLE: Design, synthesis and evaluation of substituted phenylpropanoic acid derivatives as peroxisome proliferator-activated receptor (PPAR) activators: novel human PPAR.alpha.-selective activators

AUTHOR(S): Miyachi, Hiroyuki; Nomura, Masahiro; Tanase, Takahiro; Takahashi, Yukie; Ide, Tomohiro; Tsunoda, Masaki; Murakami, Koji; Awano, Katsuya

CORPORATE SOURCE: Kyorin Pharmaceutical Co., Ltd., Discovery Research Laboratories, Tochigi, Shimotsuga-gun, Nogi-machi, 329-0114, Japan

SOURCE: Bioorganic & Medicinal Chemistry Letters (2001), Volume Date 2002, 12(1), 77-80
CODEN: BMCLE8; ISSN: 0960-894X

PUBLISHER: Elsevier Science Ltd.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB A series of substituted phenylpropanoic acid derivs. was prepd. as part of a search for subtype-selective human peroxisome proliferator-activated receptor (PPAR) activators. Structure-activity relationship studies indicated that the substituent at the .alpha.-position of the carboxyl group plays a key role in detg. the potency and the selectivity for PPAR transactivation.

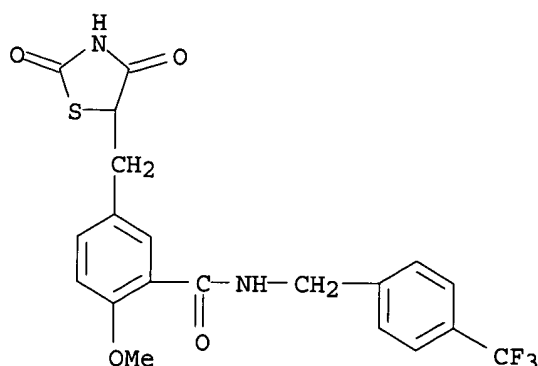
IT 213252-19-8, KRP 297

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(design, synthesis and evaluation of substituted phenylpropanoic acid derivs. as PPAR activators)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl)methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

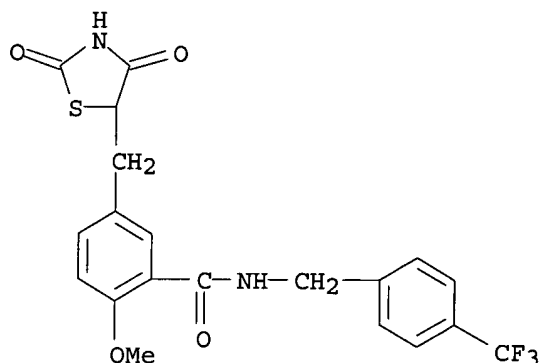
L4 ANSWER 20 OF 42 CAPLUS COPYRIGHT 2003 ACS
 ACCESSION NUMBER: 2001:798208 CAPLUS
 DOCUMENT NUMBER: 135:344474
 TITLE: Preparation of novel stable crystal of thiazolidinedione derivative
 INVENTOR(S): Oonoda, Michiro; Orita, Kazuo
 PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 17 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001081327	A1	20011101	WO 2001-JP3450	20010423
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
AU 2001048828	A5	20011107	AU 2001-48828	20010423
BR 2001010258	A	20030107	BR 2001-10258	20010423
EP 1277745	A1	20030122	EP 2001-921997	20010423
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
NO 2002005069	A	20021022	NO 2002-5069	20021022
PRIORITY APPLN. INFO.:				
			JP 2000-124006	A 20000425
			WO 2001-JP3450	W 20010423
AB	Claimed is a crystal of 5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]benzamide (KRP-297) having diffraction angles (2.theta.) at at least 9.7.degree., 15.0.degree., and 22.5.degree. in X-ray powder diffractometry. The novel crystal of KRP-297 (a known antidiabetic agent) is prepd. through recrystn. from an alc. solvent.			
IT	353275-24-8P			
	RL: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic			

preparation); PREP (Preparation); RACT (Reactant or reagent)
(prepn. of novel stable crystal of thiazolidinedione deriv.)

RN 353275-24-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]-, monosodium salt (9CI) (CA INDEX NAME)



● Na

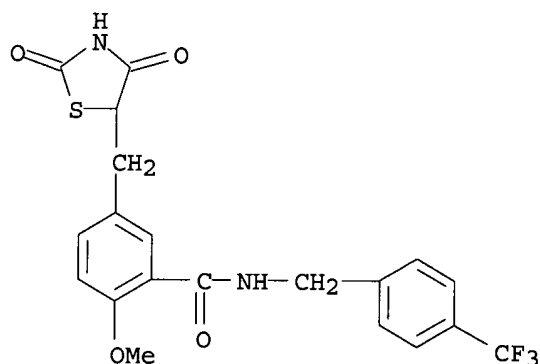
IT 213252-19-8P, KRP-297

RL: IMF (Industrial manufacture); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of novel stable crystal of thiazolidinedione deriv.)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 21 OF 42 CAPLUS COPYRIGHT 2003 ACS

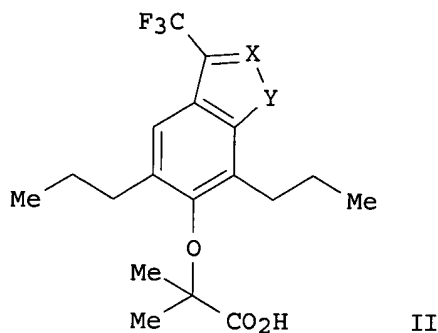
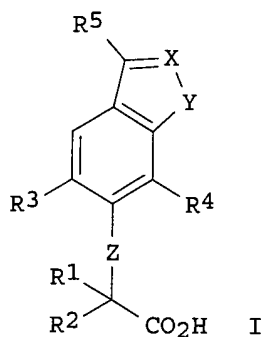
ACCESSION NUMBER: 2001:617987 CAPLUS

DOCUMENT NUMBER: 135:180757

TITLE: Preparation of 1,2-benzoxazolyloxyacetic acids and analogs as PPAR agonists for treatment of diabetes and

lipid disorders
 INVENTOR(S): Liu, Kun; Xu, Libo; Jones, A. Brian
 PATENT ASSIGNEE(S): Merck + Co. Inc., USA
 SOURCE: PCT Int. Appl., 54 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001060807	A1	20010823	WO 2001-US4636	20010214
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1259494	A1	20021127	EP 2001-910624	20010214
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
PRIORITY APPLN. INFO.:			US 2000-183593P	P 20000218
			WO 2001-US4636	W 20010214
OTHER SOURCE(S):			MARPAT 135:180757	
GI				



AB The title compds. (I) [wherein R1 and R2 = independently H, F, (halo)alkyl, (halo)alkenyl, (halo)alkynyl; or R1 and R2 may form a cycloalkyl group; R3 and R4 = independently (fluoro)alkyl, (fluoro)alkenyl, (fluoro)alkynyl, or Cl; X = N or CR; Y = O, S, nor NR; Z = O or S; R = independently H or optionally fluoro- or alkoxy-substituted (cyclo)alkyl(oxy), alkenyl(oxy), or alkynyl(oxy); R5 = H or (un)substituted alkyl, alkenyl, alkynyl, (hetero)aryl(oxy), heterocyclyl(oxy), etc.; and pharmaceutically acceptable salts and prodrugs thereof] were prepd. For example, 2,4-dihydroxy-3,5-dipropyl-1',1',1'-trifluoroacetophenone oxime was acetylated and then treated with pyridine and TEA to give 5,7-dipropyl-6-hydroxy-3-trifluoromethyl-1,2-

benzisoxazole. Etherification with Me .alpha.-bromoisobutyrate in the presence of Cs₂CO₃ in DMF, followed by sapon., afforded the 1,2-benzoxazolyloxyacetic acid (II). I are potent agonists of peroxisome proliferator activated receptor (PPAR) .alpha. and/or .gamma. and are useful in the treatment, control, or prevention of non-insulin dependent diabetes mellitus (NIDDM), hyperglycemia, dyslipidemia, hyperlipidemia, hypercholesterolemia, hypertriglyceridemia, atherosclerosis, obesity, vascular restenosis, inflammation, and other PPAR.alpha. and/or .gamma. mediated diseases, disorders, and conditions (no data).

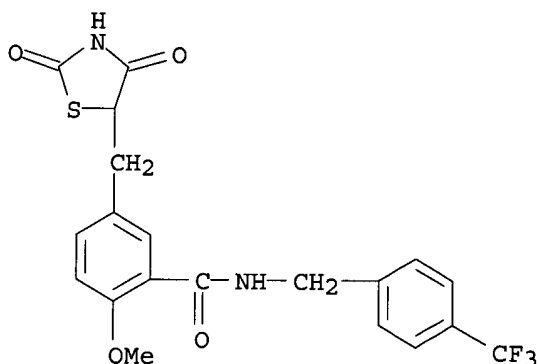
IT 213252-19-8, KRP-297

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(coadministration with; prepn. of benzisoxazolyloxyacetic acid PPAR agonists via cyclization of dihydroxyacetophenone oximes for treatment of diabetes and lipid disorders)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 22 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2001:617810 CAPLUS

DOCUMENT NUMBER: 135:175429

TITLE: Modulation of bone formation with peroxisome proliferator-activated receptor activators and ligands

INVENTOR(S): Scutt, Andrew; Still, Karen

PATENT ASSIGNEE(S): University of Sheffield, UK

SOURCE: PCT Int. Appl., 27 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

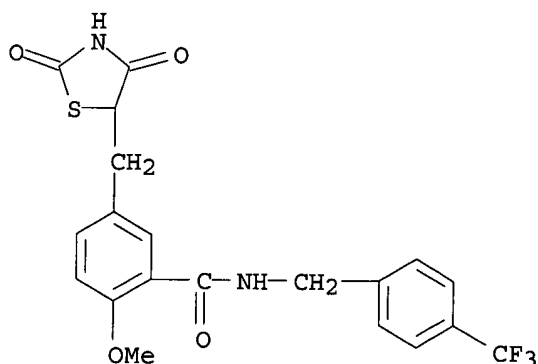
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001060355	A1	20010823	WO 2001-GB626	20010215
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,				

LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
 SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
 YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
 DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
 BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
 EP 1259233 A1 20021127 EP 2001-904207 20010215
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
 NO 2002003837 A 20021014 NO 2002-3837 20020814
 PRIORITY APPLN. INFO.: GB 2000-3310 A 20000215
 WO 2001-GB626 W 20010215
 AB The use of an activator or ligand of a peroxisome proliferator-activated
 receptor, other than PPAR.gamma., or pharmaceutically acceptable deriv. of
 said activator or ligand, in the manuf. of a medicament for the treatment
 or prophylaxis of bone disease allows, for the first time, bone anabolism
 to enhance the deposition of bone in conditions which would benefit from
 increased bone deposition. The reverse, where there is inhibition and/or
 retardation of bone deposition is also facilitated.
 IT 213252-19-8, KRP-297
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological
 study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
 (Uses)
 (modulation of bone formation with peroxisome proliferator-activated
 receptor activators and ligands)
 RN 213252-19-8 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-
 (trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS
 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 23 OF 42 CAPLUS COPYRIGHT 2003 ACS
 ACCESSION NUMBER: 2001:581862 CAPLUS
 DOCUMENT NUMBER: 135:152800
 TITLE: Alkali metal salt of thiazolidine-2,4-dione derivative
 and purification of KRP-297
 INVENTOR(S): Ohnoda, Michiro; Orita, Kazuo; Yoshida, Noriyuki
 PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan
 SOURCE: PCT Int. Appl., 17 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

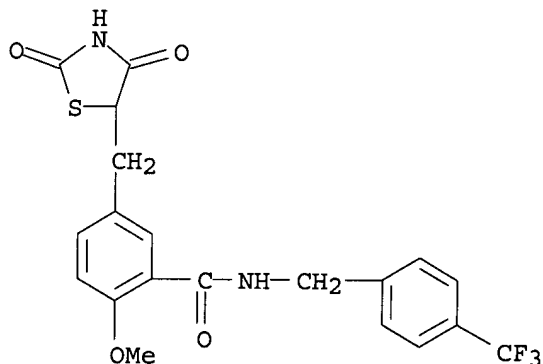
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001057007	A1	20010809	WO 2001-JP598	20010130
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
AU 2001028854	A5	20010814	AU 2001-28854	20010130
EP 1253145	A1	20021030	EP 2001-948982	20010130
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
US 2003013749	A1	20030116	US 2002-181432	20020725
PRIORITY APPLN. INFO.:				
			JP 2000-23610	A 20000201
			WO 2001-JP598	W 20010130

AB This document discloses a method of industrially advantageously purifying 5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxy-N-[4-(trifluoromethyl)phenylmethyl]benzamide (KRP-297), a known antidiabetic agent. The method comprises the steps of: forming an alkali metal salt of KRP-297 and a hydrate thereof in a reaction for producing KRP-297; isolating and purifying them; and then liberating the KRP-297 from the salt. Also provided are an alkali metal salt of KRP-297 and a hydrate of the salt.

IT **213252-19-8P**, KRP 297
 RL: PUR (Purification or recovery); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (purifn. of antidiabetic KRP-297)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)

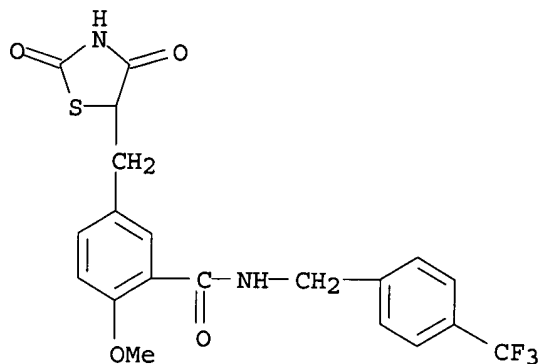


IT **353275-24-8P 353275-26-0P 353275-27-1P 353275-28-2P**
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (purifn. of antidiabetic KRP-297)

10049937

RN 353275-24-8 CAPLUS

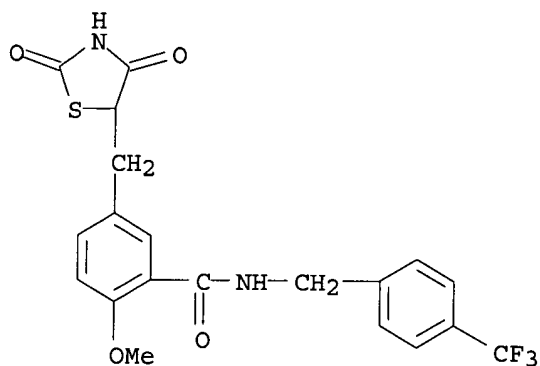
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]-, monosodium salt (9CI) (CA INDEX NAME)



● Na

RN 353275-26-0 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]-, monosodium salt, monohydrate (9CI) (CA INDEX NAME)

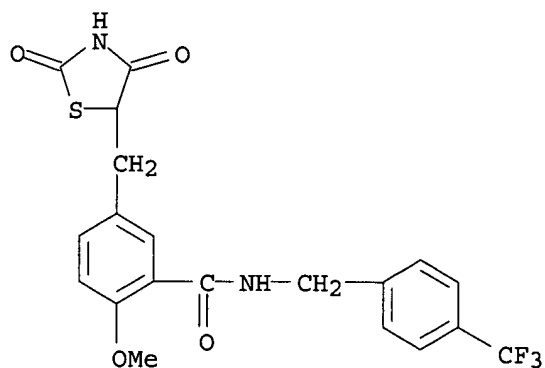


● Na

● H₂O

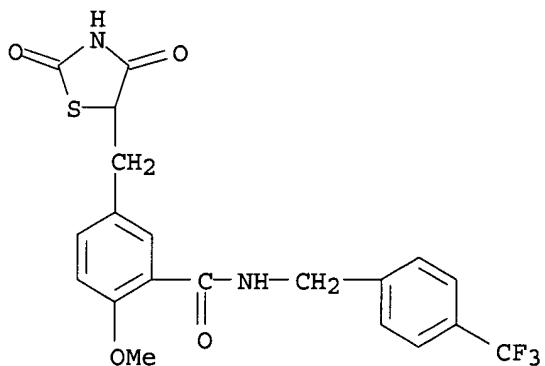
RN 353275-27-1 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]-, monopotassium salt (9CI) (CA INDEX NAME)



● K

RN 353275-28-2 CAPLUS
 CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]-, monopotassium salt, monohydrate (9CI)
 (CA INDEX NAME)



● K

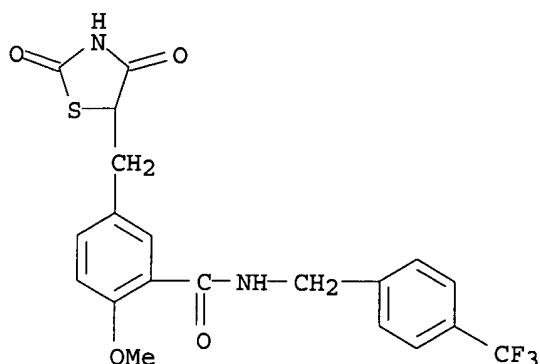
● H₂O

REFERENCE COUNT: 33 THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 24 OF 42 CAPLUS COPYRIGHT 2003 ACS
 ACCESSION NUMBER: 2001:564869 CAPLUS
 DOCUMENT NUMBER: 135:132451
 TITLE: Novel remedies with the use of .beta.3 agonists
 INVENTOR(S): Ogawa, Kohei; Umeno, Hiroshi
 PATENT ASSIGNEE(S): Asahi Kasei K. K., Japan

SOURCE: PCT Int. Appl., 49 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001054728	A1	20010802	WO 2001-JP553	20010126
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
AU 2001027103	A5	20010807	AU 2001-27103	20010126
EP 1258253	A1	20021120	EP 2001-901552	20010126
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
US 2003018061	A1	20030123	US 2002-182375	20020729
PRIORITY APPLN. INFO.:			JP 2000-20733	A 20000128
			WO 2001-JP553	W 20010126
AB	Remedies contg. at least one member selected from the group consisting of cholinolytics, monoamine reuptake inhibitors, lipase inhibitors, selective serotonin reuptake inhibitors, insulin, insulin secretion promoters, biguanide, .alpha.-glucosidase inhibitors, insulin resistance improving agents, HMC-CoA reductase inhibitors, anion exchange resins, clofibrate-base drugs and nicotinic acid-base drugs and a compd. having a .beta.3-agonistic activity. The .beta.3 agonist has an activity of inhibiting urination disorder. When used together with a remedy for urination disorder such as propiverine, oxybutynin hydrochloride or tolterodine, it exerts an enhanced anti-urination disorder effect. When used together with an antiobesity agent such as sibutramine or orlistat, it exerts an enhanced antiobesity effect. When used together with an antidiabetic agent such as insulin, glibenclamide, acarbose or rosiglitazone, it exerts an enhanced antidiabetic effect. When used together with an antilipemic drug such as bezafibrate or pravastatin, it exerts an enhanced antilipemic effect.			
IT	213252-19-8, KRP 297 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (novel remedies with the use of .beta.3 agonists as antidiabetics and antilipidemics and for treatment of urination disorder)			
RN	213252-19-8 CAPLUS			
CN	Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)			



REFERENCE COUNT: 43 THERE ARE 43 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 25 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2001:367156 CAPLUS

DOCUMENT NUMBER: 135:131731

TITLE: Design and Synthesis of 2-Methyl-2-{4-[2-(5-methyl-2-aryloxazol-4-yl)ethoxy]phenoxy}propionic Acids: A New Class of Dual PPAR.alpha./gamma. Agonists

AUTHOR(S): Brooks, Dawn A.; Etgen, Garret J.; Rito, Christopher J.; Shuker, Anthony J.; Dominianni, Samuel J.; Warshawsky, Alan M.; Ardecky, Robert; Paterniti, James R.; Tyhonas, John; Karanewsky, Donald S.; Kauffman, Raymond F.; Broderick, Carol L.; Oldham, Brian A.; Montrose-Rafizadeh, Chahzrad; Winneroski, Leonard L.; Faul, Margaret M.; McCarthy, James R.

CORPORATE SOURCE: Lilly Research Laboratories A Division of Eli Lilly Company Lilly Corporate Center, Indianapolis, IN, 46285, USA

SOURCE: Journal of Medicinal Chemistry (2001), 44(13), 2061-2064

CODEN: JMCMAR; ISSN: 0022-2623

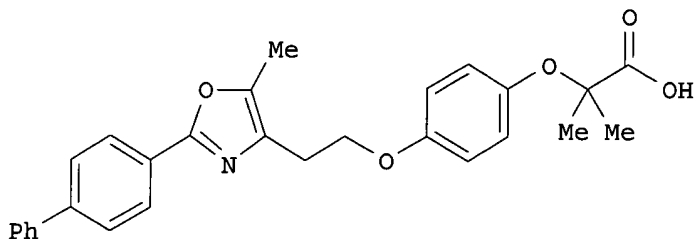
PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

OTHER SOURCE(S): CASREACT 135:131731

GI



I

AB Propionic acid deriv. I, which was designed and synthesized based on putative pharmacophores of known PPAR.gamma.- and PPAR.alpha.-selective compds., exhibits potent dual PPAR.alpha./gamma. agonist activity as

demonstrated by in vitro binding and dose overlap in the newly introduced EOB mouse model for glucose lowering and lipid/cholesterol homeostasis.

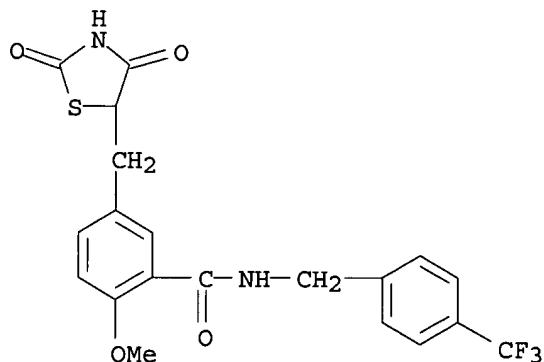
IT 213252-19-8, KRP-297

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(design and synthesis of 2-methyl-2-{4-[2-(5-methyl-2-aryloxazol-4-yl)ethoxy]phenoxy}propionic acids: a new class of dual PPAR.alpha./gamma. agonists)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 24 THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 26 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2001:359797 CAPLUS

DOCUMENT NUMBER: 134:344620

TITLE: Solid oral composition containing KRP-297

INVENTOR(S): Ohyama, Toshinori; Imamizu, Masaru

PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan

SOURCE: PCT Int. Appl., 11 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001034148	A1	20010517	WO 2000-JP7905	20001110
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
AU 2001013046	A5	20010606	AU 2001-13046	20001110
EP 1243266	A1	20020925	EP 2000-974882	20001110

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO, MK, CY, AL, TR

PRIORITY APPLN. INFO.: JP 1999-320586 A 19991111
WO 2000-JP7905 W 20001110

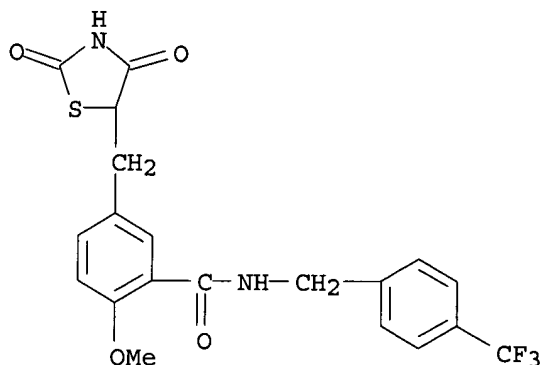
AB Disclosed are solid compns. for oral use for facilitating the administration in a small dose of KRP-297, which is a ligand common to peroxisome proliferator-activated receptors PPAR.alpha. and PPAR.gamma. (i.e., nuclear receptors) and has an effect of ameliorating insulin resistance, which contain the drug ingredient in a uniform content and can be easily and quant. taken. By prepg. solid compns. for oral use composed of a trace amt. of the drug ingredient together with pharmaceutical carriers, it is possible to provide tablets which contain the drug component in a uniform content and can be easily and quant. taken. A film-coated tablet was prepd. from KRP-297 0.25, lactose 78.55, cryst. cellulose 26.2, low-substituted hydroxypropyl cellulose 12, polyvinyl alc. 2.4, magnesium stearate 0.6, hydroxypropyl Me cellulose, and carnauba wax 0.001 mg.

IT 213252-19-8, KRP-297

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(solid oral compns. contg. uniform contents of KRP-297)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 27 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2001:347100 CAPLUS

DOCUMENT NUMBER: 134:353303

TITLE: preparation of thiazolidinyl-containing bicyclic heterocycles as humane peroxisome proliferator-activated receptor .gamma. agonists

INVENTOR(S): Nomura, Masahiro; Murakami, Koji; Kakuta, Masaki

PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 7 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent

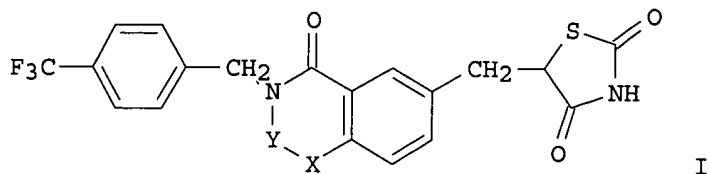
LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
------------	------	------	-----------------	------

 JP 2001131173 A2 20010515 JP 2000-242708 20000810
 PRIORITY APPLN. INFO.: JP 1999-235531 A 19990823
 OTHER SOURCE(S): MARPAT 134:353303
 GI



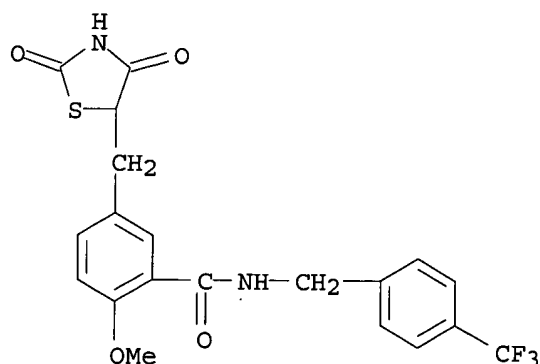
AB Title compds. I (YX = CO₂, CH₂O, CH:CH), their pharmaceutically acceptable salts, or hydrates, useful as for treatment of Type II diabetes and hyperlipemia, are prepd. 2-Hydroxy-5-[(2,4-dioxothiazolidin-5-yl)methyl]-N-[(4-trifluorophenyl)methyl]benzamide was reacted with trioxane in the presence of AcOH in CH₂Cl₂ at room temp. for 2 day to give 42% 6-[(2,4-dioxothiazolidin-5-yl)methyl]-3-[(4-trifluorophenyl)methyl]-1,3-benzoxazin-4-one showing good transcription activity of proliferator-activated receptor .gamma. in vitro.

IT **213252-19-8**

RL: RCT (Reactant); RACT (Reactant or reagent)
 (prepn. of bicyclic heterocycles as humane peroxisome
 proliferator-activated receptor .gamma. agonists)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



IT **223508-81-4P**

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(prepn. of bicyclic heterocycles as humane peroxisome
 proliferator-activated receptor .gamma. agonists)

RN 223508-81-4 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-hydroxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 28 OF 42 CAPLUS COPYRIGHT 2003 ACS
ACCESSION NUMBER: 2001:338335 CAPLUS
DOCUMENT NUMBER: 134:344604
TITLE: Antidiabetic formulation containing metformin and
sulfonylurea
INVENTOR(S): Piper, Beth Anne
PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA
SOURCE: PCT Int. Appl., 76 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 3
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001032158	A2	20010510	WO 2000-US28467	20001013
WO 2001032158	A3	20020829		
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
US 2002177602	A1	20021128	US 1999-432465	19991103
EP 1253944	A2	20021106	EP 2000-970913	20001013
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL			
NO 200202086	A	20020624	NO 2002-2086	20020502
PRIORITY APPLN. INFO.:			US 1999-432465	A 19991103
			WO 2000-US28467	W 20001013

AB A low dose antidiabetic formulation adapted for treating e.g., Type II diabetes contains a combination of metformin (at <800 mg/day) and at least 1 other antidiabetic agent such as a sulfonylurea. This combination provides at least about substantially equiv. efficacy in treating diabetes as do antidiabetic formulations contg. metformin employed in dosages prescribed in generally accepted medical practice for first line therapy in treating diabetes, but with substantially reduced side effects, such as hypoglycemia and/or gastrointestinal distress. A method for treating diabetes in drug naive human patients is also provided employing the above

formulation to reduce insulin resistance and/or post-prandial glucose excursion and/or Hb 1Ac, and/or increase post-prandial insulin, thereby treating the diabetes. Thus, tablets contained metformin-HCl 250.0, glyburide 1.25, croscarmellose sodium 7.00, Povidone 10.00, microcryst. cellulose 28.25, Mg stearate 2.25, and HPMC film-coating 6 mg. The effectiveness of this combination drug in producing hypoglycemia was demonstrated clin.

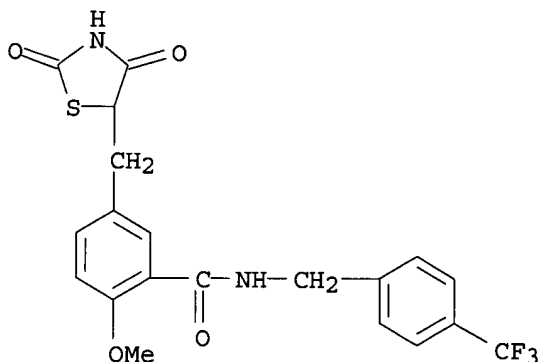
IT 213252-19-8, KRP-297

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(antidiabetic formulation contg. metformin and sulfonylurea)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 29 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2001:283949 CAPLUS

DOCUMENT NUMBER: 134:311218

TITLE: Synthesis and use of heterocyclic sodium/proton exchange inhibitors

INVENTOR(S): Ahmad, Saleem; Wu, Shung C.; O'Neil, Steven V.; Ngu, Khehyong; Atwal, Karnail S.

PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA

SOURCE: PCT Int. Appl., 221 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

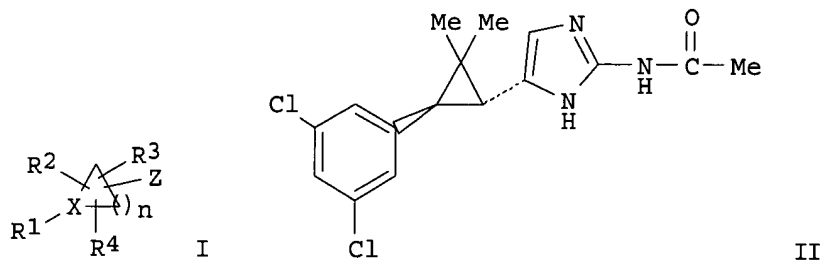
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001027107	A2	20010419	WO 2000-US27461	20001002
WO 2001027107	A3	20020124		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,				

DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
 CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
 EP 1224183 A2 20020724 EP 2000-968723 20001002
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO, MK, CY, AL
 NO 2002001717 A 20020610 NO 2002-1717 20020411
 PRIORITY APPLN. INFO.: US 1999-158755P P 19991012
 WO 2000-US27461 W 20001002
 OTHER SOURCE(S): MARPAT 134:311218
 GI



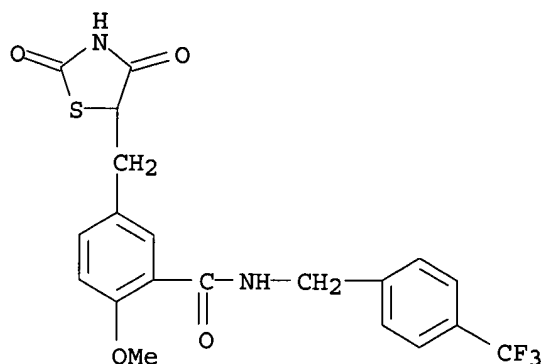
AB Compds. of formula I [wherein; n is 1-5; X is N or CR₅, where R₅ is H, halo, alkenyl, alkynyl, alkoxy, alkyl, aryl or heteroaryl; Z is a heteroaryl group; R₁ is H, alk(en)(yn)yl, alk(enyl)(ynyl)oxy, (aryl or alkyl)3Si, cycloalk(en)yl, (aryl)amino, aryl(alkyl), cycloheteroaryl, etc.; R₂, R₃ and R₄ are any of the groups set out for R₁ and optionally substituted with 1 to 5 substituents which may be the same or different and when X is N, R₁ is preferably aryl or heteroaryl] are claimed. Several hundred examples are disclosed. Synthesis of II proceeds via cyclopropanation of the cinnamate derived from the olefination between 3,5-dichlorobenzaldehyde and t-butyldiethylphosphonoacetate. The intermediate tert-Bu ester is converted to the corresponding .alpha.-chloroketone and reacted with acetyl guanidine to provide II in a total of 5 steps. Compds. I are said to be sodium/proton exchange inhibitors (NHE). Pharmaceutical combinations are claimed using I and certain antihypertensive agents, .beta.-adrenergic agonists, hypolipidemic agents, antidiabetic agents, antiobesity agents, etc. Compds. I are useful as antianginal and cardioprotective agents and provide a method for preventing or treating angina pectoris, cardiac dysfunction, myocardial necrosis, and arrhythmia.

IT 213252-19-8, KRP297
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pharmaceuticals also contg.; synthesis and use of heterocyclic sodium/proton exchange inhibitors)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 30 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2001:152661 CAPLUS

DOCUMENT NUMBER: 134:193428

TITLE: Preparation of substituted benzylthiazolidine-2,4-dione derivatives as agonists of human peroxisome proliferator-activated receptor

INVENTOR(S): Nomura, Masahiro; Murakami, Koji; Tsunoda, Masaki; Takahashi, Yukie

PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan

SOURCE: PCT Int. Appl., 19 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

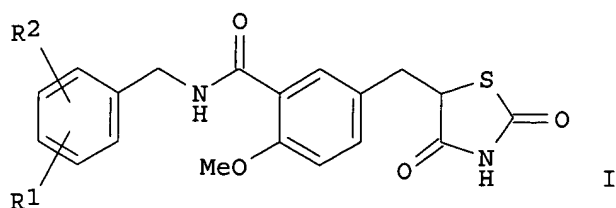
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001014352	A1	20010301	WO 2000-JP5522	20000818
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1207158	A1	20020522	EP 2000-953478	20000818
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL				

PRIORITY APPLN. INFO.: JP 1999-235530 A 19990823
WO 2000-JP5522 W 20000818

OTHER SOURCE(S): MARPAT 134:193428

GI

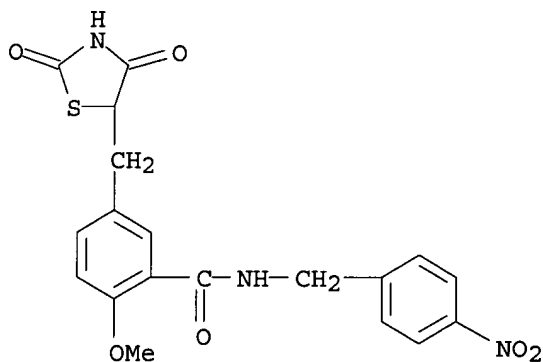


AB The title compds. (I), pharmaceutically acceptable salts thereof and hydrates of the same (wherein R1 represents chloro, bromo, nitro, trifluoromethoxy, ethoxy, propoxy or isopropoxy; and R2 represents hydrogen or chloro) are prepd. These compds. are capable of, as a ligand of human peroxisome proliferator-activated receptor (PPAR), enhancing the transcriptional activity of the receptor and showing effects of lowering blood sugar level and lowering lipid level; and a process for producing the same. Thus, 5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxybenzoic acid, Et3N, and CH2Cl2 were mixed, treated with Et chlorocarbonate and stirred under ice-cooling for 10 min, treated with 4-nitrobenzylamine, and then stirred at room temp. for 2 h to give 75% N-[(4-nitrophenyl)methyl]-5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxybenzamide (II). II and I (R1 = 4-n-Pro, R2 = H) enhanced the transcriptional activity of human PPAR.alpha. in CHO cells with EC50 of 0.53 and 0.11 .mu.M, resp.

IT 326926-46-9P 326926-47-0P 326926-48-1P
 326926-49-2P 326926-50-5P 326926-51-6P
 326926-52-7P 326926-53-8P 326926-54-9P,
 N-[(3,4-Dichlorophenyl)methyl]-5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxybenzamide
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (prepn. of substituted benzylthiazolidinedione derivs. as agonists of human peroxisome proliferator-activated receptor and blood sugar and lipid-lowering agents)

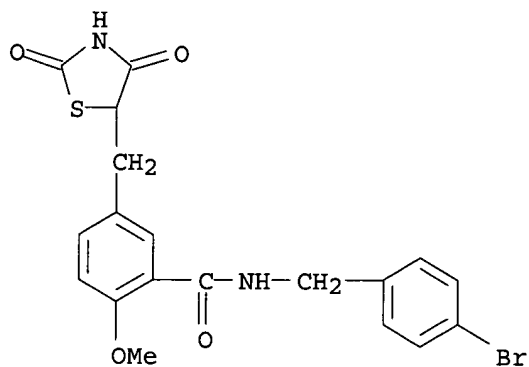
RN 326926-46-9 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-nitrophenyl)methyl]- (9CI) (CA INDEX NAME)



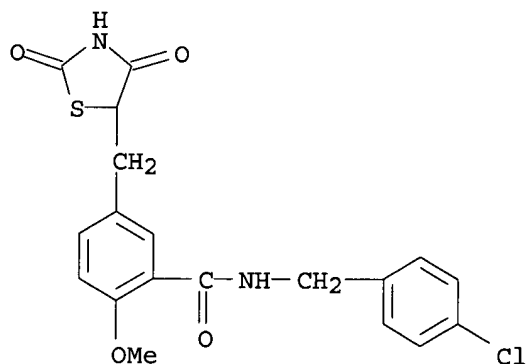
RN 326926-47-0 CAPLUS

CN Benzamide, N-[(4-bromophenyl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



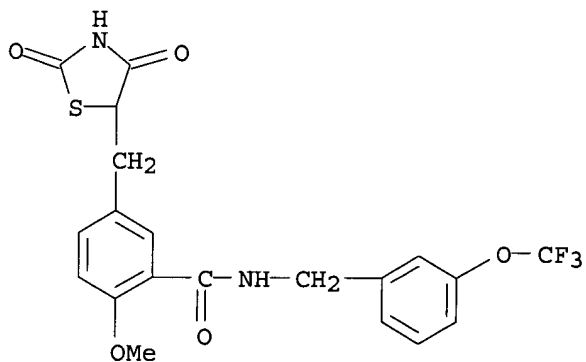
RN 326926-48-1 CAPLUS

CN Benzamide, N-[(4-chlorophenyl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



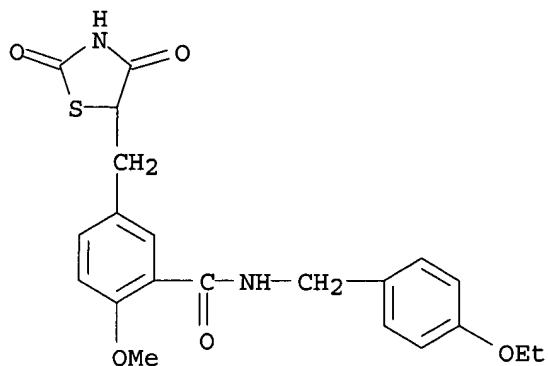
RN 326926-49-2 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(3-(trifluoromethoxy)phenyl)methyl]- (9CI) (CA INDEX NAME)



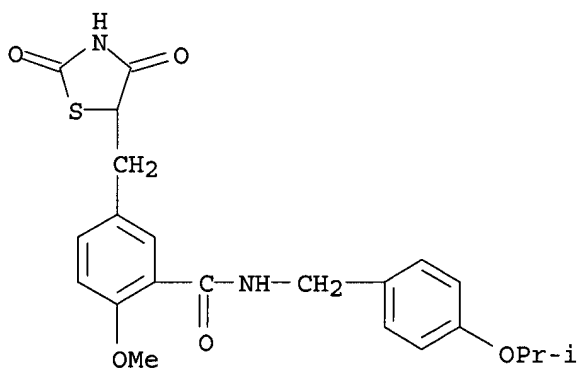
RN 326926-50-5 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-N-[(4-ethoxyphenyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



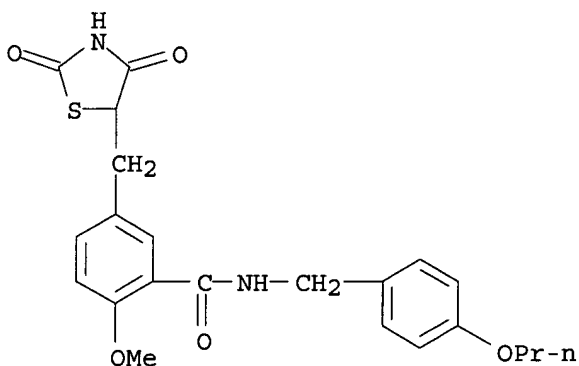
RN 326926-51-6 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(1-methylethoxy)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 326926-52-7 CAPLUS

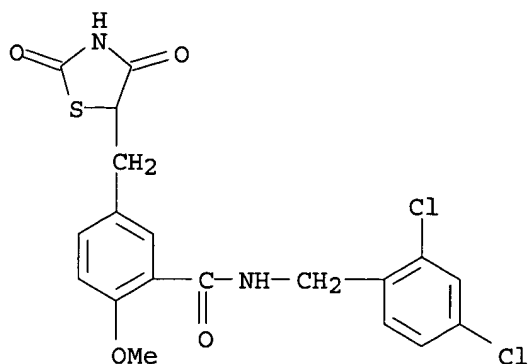
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-propoxyphenyl)methyl]- (9CI) (CA INDEX NAME)



RN 326926-53-8 CAPLUS

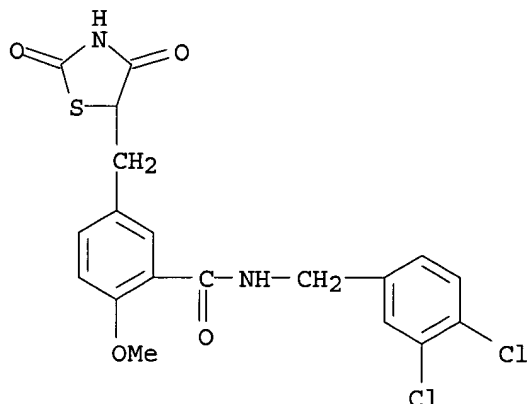
CN Benzamide, N-[(2,4-dichlorophenyl)methyl]-5-[(2,4-dioxo-5-

thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



RN 326926-54-9 CAPLUS

CN Benzamide, N-[(3,4-dichlorophenyl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 31 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2001:152660 CAPLUS

DOCUMENT NUMBER: 134:193427

TITLE: Preparation of substituted benzylthiazolidine-2,4-dione derivatives as agonists of human peroxisome proliferator-activated receptor

INVENTOR(S): Miyachi, Hiroyuki; Nomura, Masahiro; Tanase, Takahiro; Murakami, Koji; Tsunoda, Masaki

PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan

SOURCE: PCT Int. Appl., 20 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

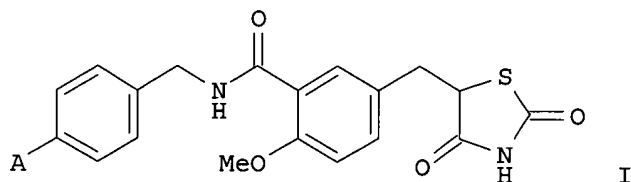
PATENT INFORMATION:

PATENT NO.

KIND DATE

APPLICATION NO. DATE

 WO 2001014351 A1 20010301 WO 2000-JP5521 20000818
 W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU,
 CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL,
 IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD,
 MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK,
 SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY,
 KG, KZ, MD, RU, TJ, TM
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
 DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
 CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
 EP 1207157 A1 20020522 EP 2000-953477 20000818
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO, MK, CY, AL
 PRIORITY APPLN. INFO.: JP 1999-235529 A 19990823
 JP 2000-242707 A 20000810
 WO 2000-JP5521 W 20000818
 OTHER SOURCE(S): MARPAT 134:193427
 GI



AB The title compds. represented by general formula (I; wherein A represents optionally substituted Ph, optionally substituted phenoxy or optionally substituted benzyloxy), pharmaceutically acceptable salts thereof and hydrates of the same are prepd. These compds. are capable of, as a ligand of human peroxisome proliferator-activated receptor (PPAR), enhancing the transcriptional activity of the receptor and showing effects of lowering blood sugar level and lowering lipid level. Thus, 5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxybenzoic acid, Et₃N, and CH₂Cl₂ were mixed, treated with Et chlorocarbonate under ice-cooling, and stirred for 10 min under ice-cooling, followed by adding a soln. of 4-benzyloxybenzylamine in CH₂Cl₂, and the resulting mixt. was stirred at room temp. for 2 h to give 77% N-[(4-benzyloxyphenyl)methyl]-5-[(2,4-dioxothiazolidin-5-yl)methyl]-2-methoxybenzamide (II). II and I (A = PhO) enhanced the transcriptional activity of human PPAR.alpha. in CHO cells with EC₅₀ of 0.44 and 0.24 .mu.M, resp.

IT 326925-77-3P 326925-78-4P 326925-79-5P
 326925-80-8P 326925-81-9P 326925-82-0P
 326925-83-1P 326925-84-2P 326925-85-3P
 326925-86-4P 326925-87-5P 326925-88-6P
 326925-89-7P

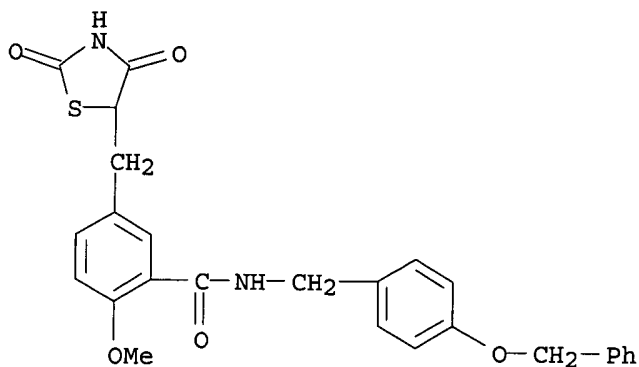
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of substituted benzylthiazolidinedione derivs. as agonists of human peroxisome proliferator-activated receptor and blood sugar and lipid-lowering agents)

RN 326925-77-3 CAPLUS

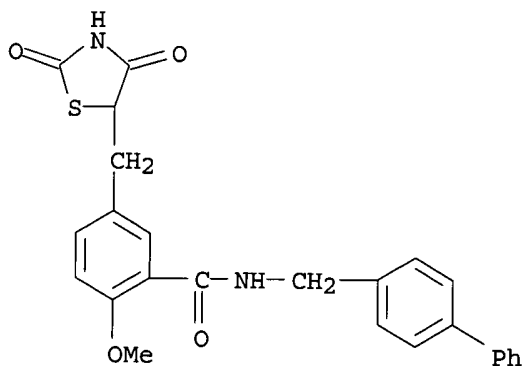
10049937

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(phenylmethoxy)phenyl]methyl]- (9CI) (CA INDEX NAME)



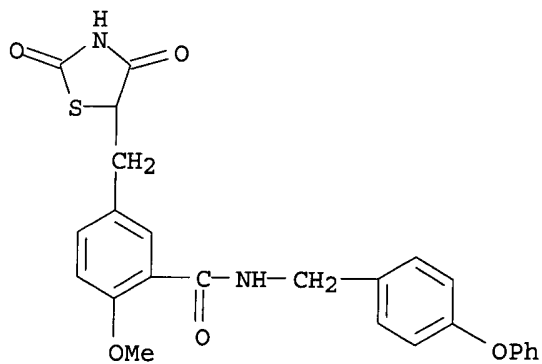
RN 326925-78-4 CAPLUS

CN Benzamide, N-([1,1'-biphenyl]-4-ylmethyl)-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



RN 326925-79-5 CAPLUS

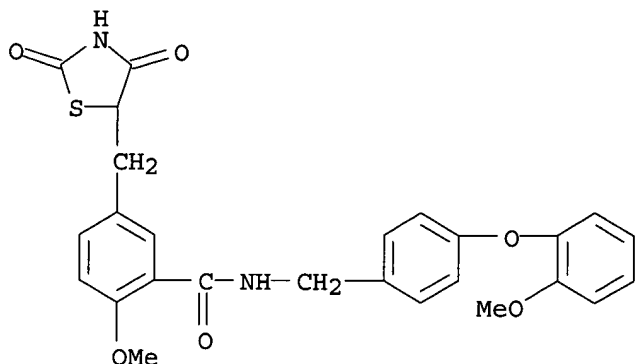
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-phenoxyphenyl)methyl]- (9CI) (CA INDEX NAME)



10049937

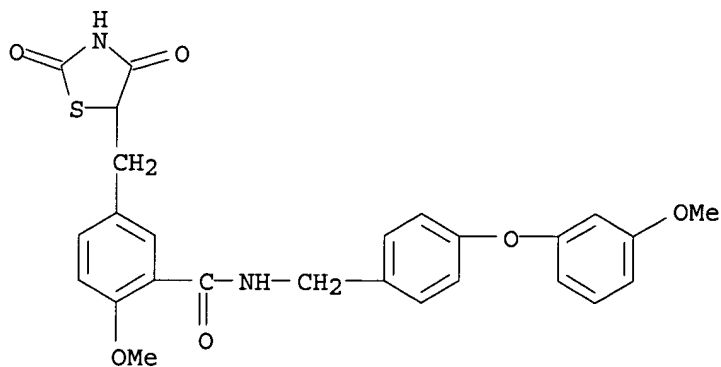
RN 326925-80-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(2-methoxyphenoxy)phenyl]methyl]- (9CI) (CA INDEX NAME)



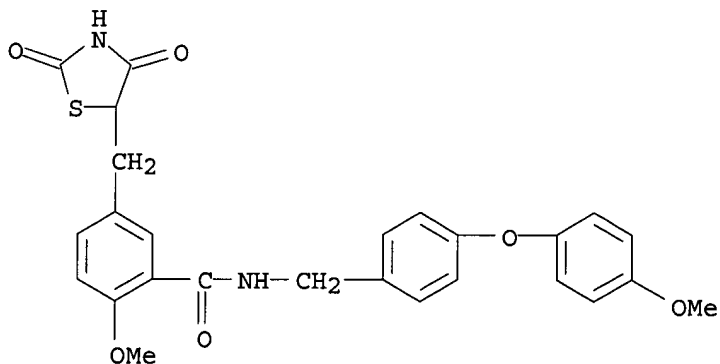
RN 326925-81-9 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(3-methoxyphenoxy)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 326925-82-0 CAPLUS

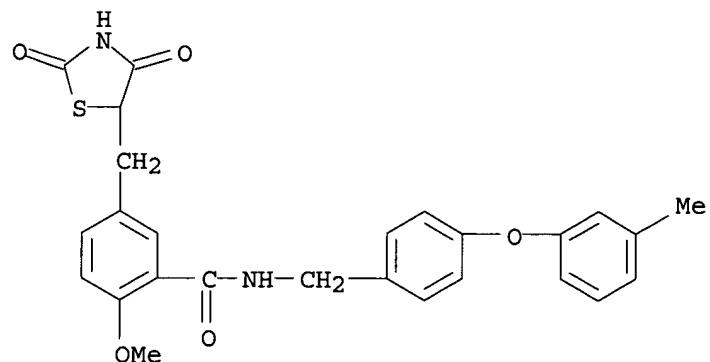
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(4-methoxyphenoxy)phenyl]methyl]- (9CI) (CA INDEX NAME)



10049937

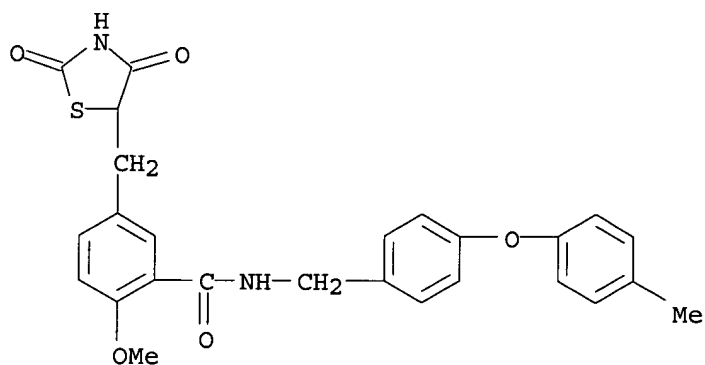
RN 326925-83-1 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(3-methylphenoxy)phenyl]methyl]- (9CI) (CA INDEX NAME)



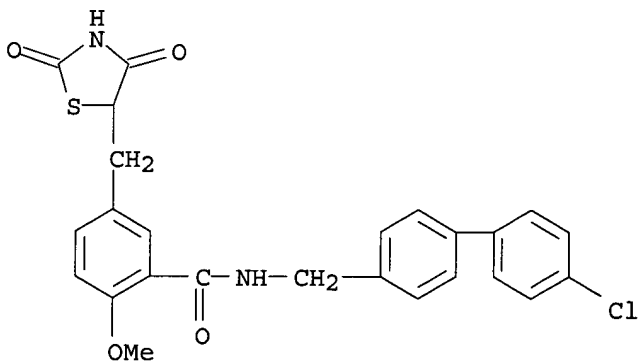
RN 326925-84-2 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(4-methylphenoxy)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 326925-85-3 CAPLUS

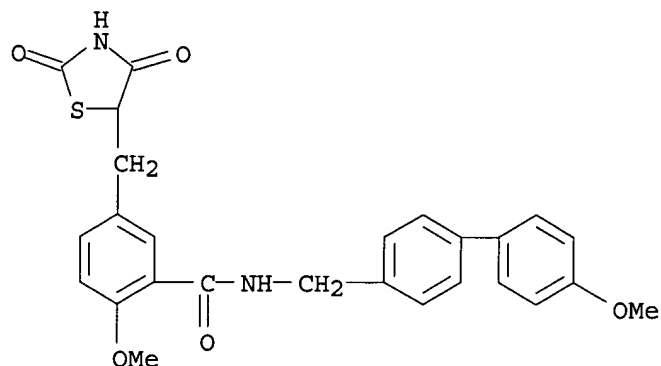
CN Benzamide, N-[(4'-chloro[1,1'-biphenyl]-4-yl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



10049937

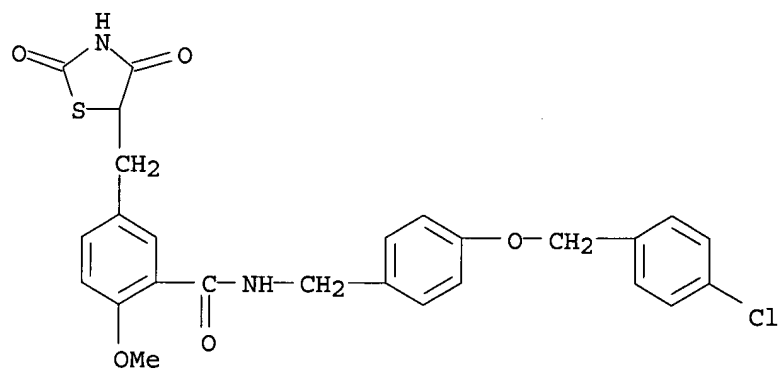
RN 326925-86-4 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4'-methoxy[1,1'-biphenyl]-4-yl)methyl]- (9CI) (CA INDEX NAME)



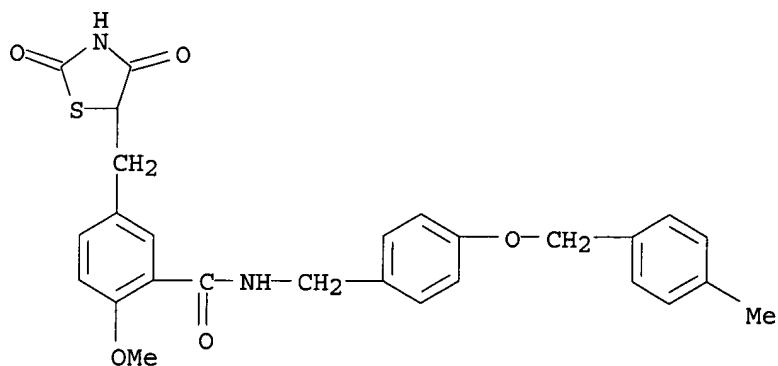
RN 326925-87-5 CAPLUS

CN Benzamide, N-[[4-[(4-chlorophenyl)methoxy]phenyl]methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



RN 326925-88-6 CAPLUS

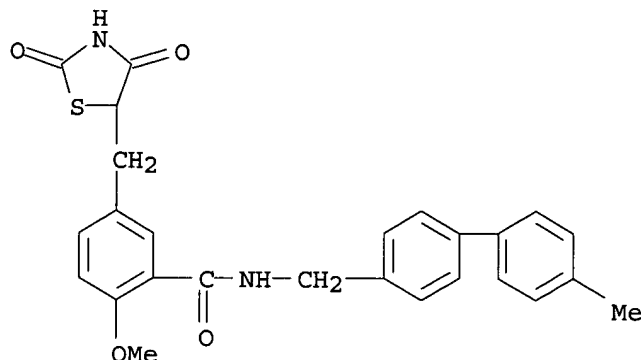
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-[(4-methylphenyl)methoxy]phenyl]methyl]- (9CI) (CA INDEX NAME)



10049937

RN 326925-89-7 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4'-methyl[1,1'-biphenyl]-4-yl)methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 32 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2000:293502 CAPLUS

DOCUMENT NUMBER: 133:84110

TITLE: Fenofibrate and Rosiglitazone Lower Serum

Triglycerides with Opposing Effects on Body Weight

AUTHOR(S): Chaput, Evelyne; Saladin, Regis; Silvestre, Martine; Edgar, Alan D.

CORPORATE SOURCE: Department of Metabolic Diseases, Laboratoire Fournier, Daix, 21121, Fr.

SOURCE: Biochemical and Biophysical Research Communications (2000), 271(2), 445-450

CODEN: BBRCA9; ISSN: 0006-291X

PUBLISHER: Academic Press

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Activators of peroxisome proliferator activated receptors (PPARs) are effective drugs to improve the metabolic abnormalities linking hypertriglyceridemia to diabetes, hyperglycemia, insulin-resistance, and atherosclerosis. We compared the pharmacol. profile of a PPAR.alpha. activator, fenofibrate, and a PPAR.gamma. activator, rosiglitazone, on serum parameters, target gene expression, and body wt. gain in (fa/fa) fatty Zucker rats and db/db mice as well as their assocn. in db/db mice. Fenofibrate faithfully modified the expression of PPAR.alpha. responsive genes. Rosiglitazone increased adipose tissue aP2 mRNA in both models while increasing liver acyl CoA oxidase mRNA in db/db mice but not in fatty Zucker rats. Both drugs lowered serum triglycerides yet rosiglitazone markedly increased body wt. gain while fenofibrate decreased body wt. gain in fatty Zucker rats. KRP 297, which has been reported to be a PPAR.alpha. and .gamma. co-activator, also affected serum triglycerides and insulin in fatty Zucker rats although no change in body wt. gain was noted. These results serve to clearly differentiate the metabolic finality of two distinct classes of drugs, as well as their corresponding nuclear receptors, having similar effects on serum triglycerides. (c) 2000 Academic Press.

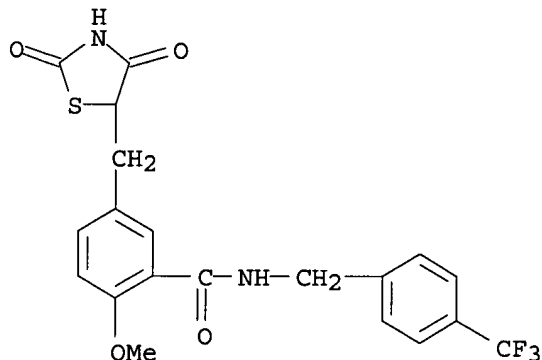
IT 213252-19-8, KRP 297

RL: BAC (Biological activity or effector, except adverse); BSU (Biological

study, unclassified); BIOL (Biological study)
 (fenofibrate and rosiglitazone lower serum triglycerides with opposing
 effects on body wt.)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 33 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2000:243901 CAPLUS

DOCUMENT NUMBER: 133:12622

TITLE: Tissue-specific actions of antidiabetic thiazolidinediones on the reduced fatty acid oxidation in skeletal muscle and liver of Zucker diabetic fatty rats

AUTHOR(S): Ide, Tomohiro; Nakazawa, Tomoko; Mochizuki, Toshiro; Murakami, Koji

CORPORATE SOURCE: Central Research Laboratories, Kyorin Pharmaceutical, Tochigi, 329-0114, Japan

SOURCE: Metabolism, Clinical and Experimental (2000), 49(4), 521-525

CODEN: METAAJ; ISSN: 0026-0495

PUBLISHER: W. B. Saunders Co.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Fatty acid overload has been proposed as a cause of decreased responsiveness in the major insulin target tissues of the body such as muscle and liver tissue. We therefore investigated fatty acid oxidn. in soleus muscle and liver isolated from Zucker diabetic fatty (ZDF) rats treated with thiazolidinediones, a new class of antidiabetic agents. ¹⁴C¹⁴O₂ prodn. from [¹⁴C]palmitic (C16:0) acid was lower in the soleus muscle and liver of ZDF rats vs. lean rats (P < .05). When administered orally to ZDF rats for 2 wk, the thiazolidinediones troglitazone (300 mg/kg) and KRP-297 (10 mg/kg) increased palmitic acid oxidn. in the soleus muscle of ZDF rats (P < .05). KRP-297, but not troglitazone, increased palmitic acid oxidn. in the liver of ZDF rats (P < .05), and both troglitazone and KRP-297 inhibited triglyceride accumulation in the skeletal muscle of ZDF rats. Hepatic triglyceride accumulation in ZDF rats was inhibited by KRP-297, but not by troglitazone. A redn. of fatty acid oxidn. in the liver of ZDF rats and an increase in response to KRP-297 were obsd. only when C16:0 and C18:0 fatty acids, not C8:0, were

used as substrates. Thus, there were defects in fatty acid catabolic activity and triglyceride accumulation in the soleus muscle and liver of ZDF rats. These results indicate that KRP-297 has advantages over troglitazone in the amelioration of these lipid metabolic abnormalities in insulin resistance assocd. with obesity.

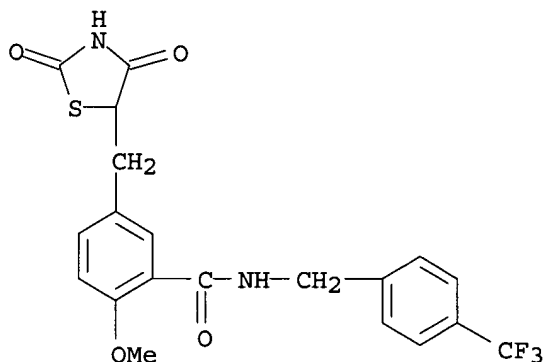
IT 213252-19-8, KRP-297

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(tissue-specific actions of antidiabetic thiazolidinediones on reduced fatty acid oxidn. in muscle and liver in NIDDM/obesity)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 34 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2000:190928 CAPLUS

DOCUMENT NUMBER: 132:231969

TITLE: Method for treating diabetes employing an ap2 inhibitor and combination

INVENTOR(S): Robl, Jeffrey A.; Parker, Rex A.; Biller, Scott A.; Jamil, Haris; Jacobson, Bruce L.; Kodukula, Krishna

PATENT ASSIGNEE(S): Bristol-Myers Squibb Co., USA

SOURCE: PCT Int. Appl., 55 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000015229	A1	20000323	WO 1999-US20946	19990913
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,				

ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
 CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

CA 2344309	AA	20000323	CA 1999-2344309	19990913
AU 9963877	A1	20000403	AU 1999-63877	19990913
AU 754488	B2	20021114		
BR 9913833	A	20010529	BR 1999-13833	19990913
EP 1121129	A1	20010808	EP 1999-951438	19990913

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO

EE 200100154	A	20021216	EE 2001-200100154	19990913
NO 2001001351	A	20010511	NO 2001-1351	20010316
LT 4871	B	20011227	LT 2001-22	20010316
LT 4870	B	20011227	LT 2001-23	20010316
LV 12686	B	20011020	LV 2001-57	20010412
US 2002035064	A1	20020321	US 2001-905235	20010713

PRIORITY APPLN. INFO.: US 1998-100677P P 19980917
 US 1999-390275 B1 19990907
 WO 1999-US20946 W 19990913

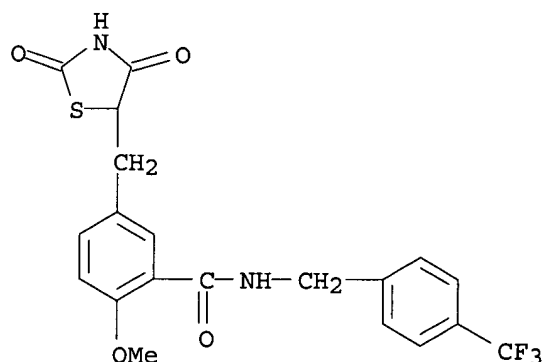
OTHER SOURCE(S): MARPAT 132:231969

AB A method is provided for treating diabetes and related diseases, such as insulin resistance, obesity, hyperglycemia, hyperinsulinemia, elevated blood levels of free fatty acids or glycerol, hypertriglyceridemia, and esp. Type II diabetes, employing an adipocyte protein aP2 inhibitor or a combination of an aP2 inhibitor and another antidiabetic agent such as metformin, glyburide, troglitazone and/or insulin.

IT **213252-19-8**, KRP 297
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (aP2 inhibitor and combination with another antidiabetic agent for treatment of diabetes and related diseases)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 35 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1999:751167 CAPLUS

DOCUMENT NUMBER: 132:44794

TITLE: Amelioration by KRP-297, a new thiazolidinedione, of impaired glucose uptake in skeletal muscle from obese

insulin-resistant animals
 AUTHOR(S): Murakami, Koji; Tsunoda, Masaki; Ide, Tomohiro;
 Ohashi, Mitsuo; Mochizuki, Toshiro
 CORPORATE SOURCE: Central Research Laboratories, Kyorin Pharmaceutical
 Co Ltd., Tochigi, Japan
 SOURCE: Metabolism, Clinical and Experimental (1999), 48(11),
 1450-1454
 CODEN: METAAJ; ISSN: 0026-0495
 PUBLISHER: W. B. Saunders Co.
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB We examd. the effect of KRP-297, a new thiazolidinedione deriv., on
 glucose uptake in the soleus muscle of two animal models of insulin
 resistance that show moderate (ob/ob mice) and severe (db/db mice)
 hyperglycemia. Insulin-stimulated 2-deoxyglucose (2DG) uptake in soleus
 muscle was 53.8% lower in ob/ob mice vs. lean mice ($P < .05$). When
 administered to ob/ob mice, KRP-297 (0.3 to 10 mg/kg) decreased plasma
 glucose and insulin levels and improved the impaired insulin-stimulated
 2DG uptake in soleus muscle in a dose-dependent manner. Soleus muscle
 from db/db mice exhibited defects in both basal (35.0% decrease, $P < .01$)
 and insulin-stimulated (50.5% decrease, $P < .01$) 2DG uptake. These
 defects were improved by treatment with KRP-297 (0.3 to 10 mg/kg).
 Moreover, KRP-297 prevented severe hyperglycemia and the marked decrease
 in pancreatic insulin content in db/db mice. These results suggest that
 KRP-297 treatment is useful to prevent the development of diabetic
 syndromes in addn. to ameliorating the impaired glucose transport in
 skeletal muscle.

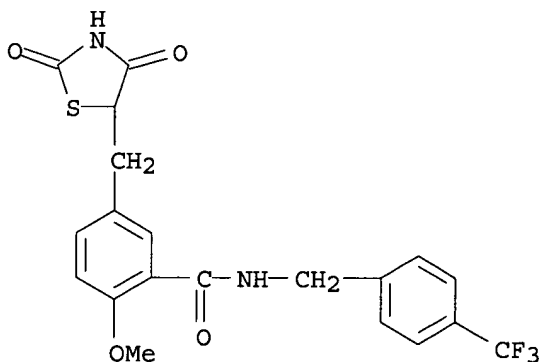
IT 213252-19-8, KRP-297

RL: BAC (Biological activity or effector, except adverse); BSU (Biological
 study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
 (Uses)

(thiazolidinedione deriv. KRP-297 amelioration of impaired glucose
 uptake in skeletal muscle from obese insulin-resistant animals)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-
 (trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS
 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 36 OF 42 CAPLUS COPYRIGHT 2003 ACS
 ACCESSION NUMBER: 1999:436161 CAPLUS
 DOCUMENT NUMBER: 131:238315

TITLE: Evidence for direct binding of fatty acids and eicosanoids to human peroxisome proliferator-activated receptor .alpha.

AUTHOR(S): Murakami, Koji; Ide, Tomohiro; Suzuki, Masahiro; Mochizuki, Toshiro; Kadowaki, Takashi

CORPORATE SOURCE: Central Research Laboratories, Kyorin Pharmaceutical Co., Ltd., Tochigi, Japan

SOURCE: Biochemical and Biophysical Research Communications (1999), 260(3), 609-613
CODEN: BBRC A9; ISSN: 0006-291X

PUBLISHER: Academic Press

DOCUMENT TYPE: Journal

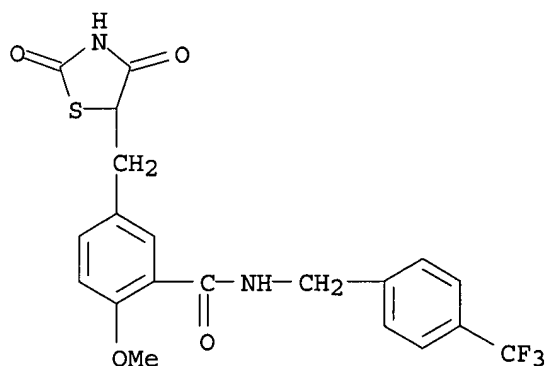
LANGUAGE: English

AB The .alpha. isoform of peroxisome proliferator-activated receptor (PPAR) is activated by fatty acids, their metabolites, and the fibrate class of lipid-lowering agents. To test the ability of these activators to directly bind the ligand-binding domain of human PPAR.alpha., we performed a competitive binding assay using radiolabeled [3H]KRP-297, a known ligand for human PPAR.alpha.. Long-chain fatty acids and eicosanoids were even more potent ligands for human PPAR.alpha. than the hitherto most potent PPAR.alpha. ligand WY-14,643. Moreover, these natural ligands avidly activated this receptor in a transient transcriptional assay. This study provides the direct evidence that human PPAR.alpha. is activated through the direct binding of fatty acids and eicosanoids, as well as of a fibrate, to its ligand-binding domain. (c) 1999 Academic Press.

IT 213252-19-8, KRP-297
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)
(direct binding of fatty acids and eicosanoids to human peroxisome proliferator-activated receptor .alpha.)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 24 THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

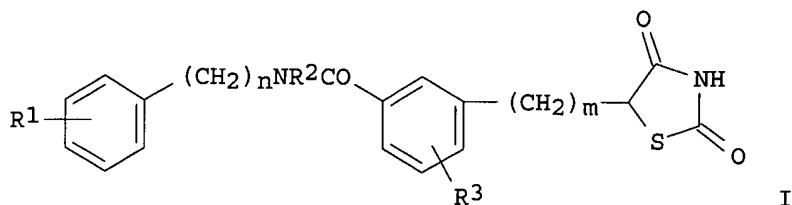
L4 ANSWER 37 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1999:188591 CAPLUS

DOCUMENT NUMBER: 130:311725

TITLE: (3-Substituted benzyl)thiazolidine-2,4-diones as structurally new antihyperglycemic agents

AUTHOR(S): Nomura, Masahiro; Kinoshita, Susumu; Satoh, Hiroya;
 Maeda, Toshio; Murakami, Koji; Tsunoda, Masaki;
 Miyachi, Hiroyuki; Awano, Katsuya
 CORPORATE SOURCE: Central Research Laboratories, Kyorin Pharmaceutical
 Co., Ltd., Tochigi, 329-0114, Japan
 SOURCE: Bioorganic & Medicinal Chemistry Letters (1999), 9(4),
 533-538
 CODEN: BMCLE8; ISSN: 0960-894X
 PUBLISHER: Elsevier Science Ltd.
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 GI



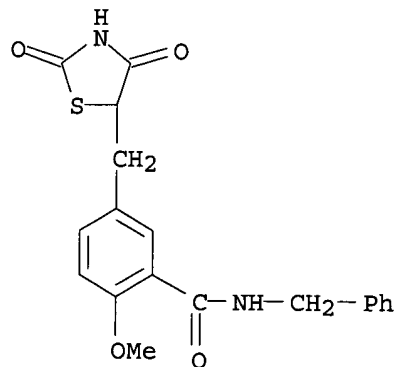
AB Title compds. I (R_1 = 4-tert-Bu, H, 4-Me, 4-MeO, 4-CF₃, etc.; R_2 = H, Et; R_3 = 6-MeO, 4-MeO, 2-MeO, 6-EtO, 6-OH, 6-F, etc.; m = 0-3; n = 0-2) were prepd. A structure-activity study of these compds. led to the identification of I (R_1 = CF₃, R_2 = H, R_3 = 6-MeO, m = n = 1) (KRP-297) as a candidate drug for the treatment of diabetes mellitus.

IT 185808-38-2P 185808-40-6P 185808-42-8P
 185808-45-1P 185808-49-5P 185808-51-9P
 185808-52-0P 185808-55-3P 185808-59-7P
 185808-62-2P 185808-63-3P 185808-64-4P
 185808-65-5P 185808-67-7P 185808-68-8P
 185808-70-2P 186312-86-7P 213252-19-8P,
 KRP-297 223508-81-4P 223508-82-5P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)
 (prepn. and antihyperglycemic activity of)

RN 185808-38-2 CAPLUS

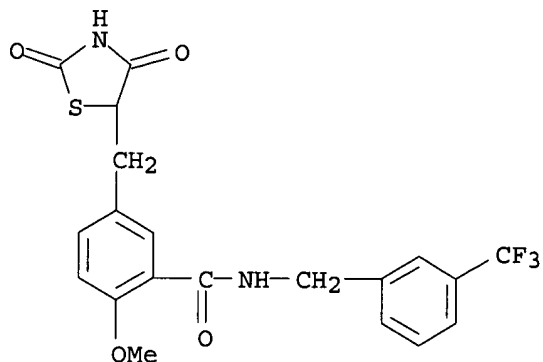
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-(phenylmethyl)- (9CI) (CA INDEX NAME)



10049937

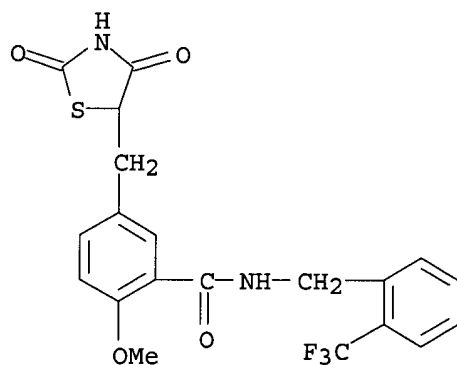
RN 185808-40-6 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[3-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



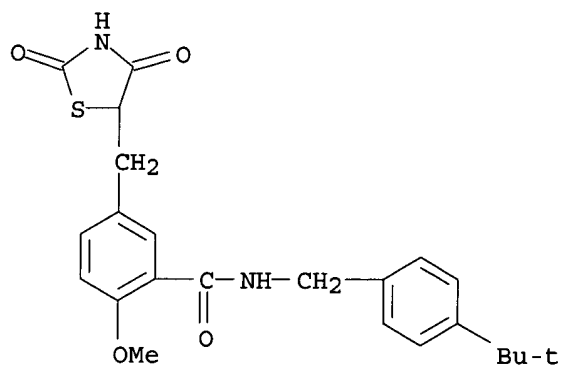
RN 185808-42-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[2-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 185808-45-1 CAPLUS

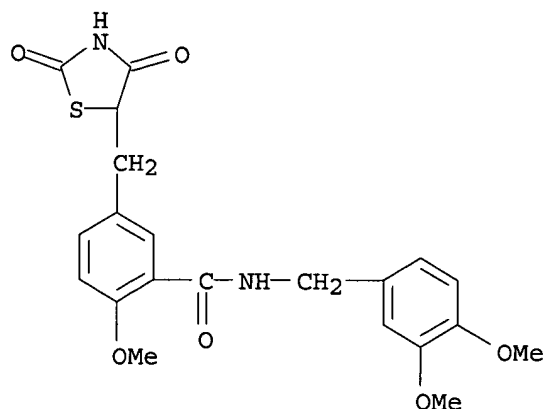
CN Benzamide, N-[[4-(1,1-dimethylethyl)phenyl]methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



10049937

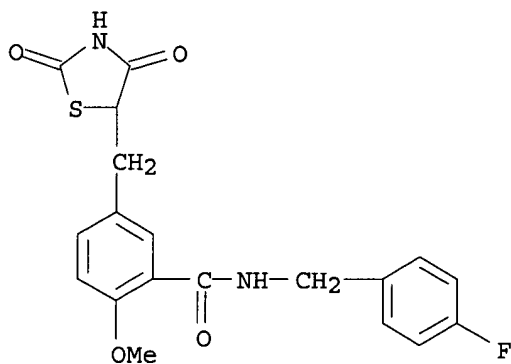
RN 185808-49-5 CAPLUS

CN Benzamide, N-[(3,4-dimethoxyphenyl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



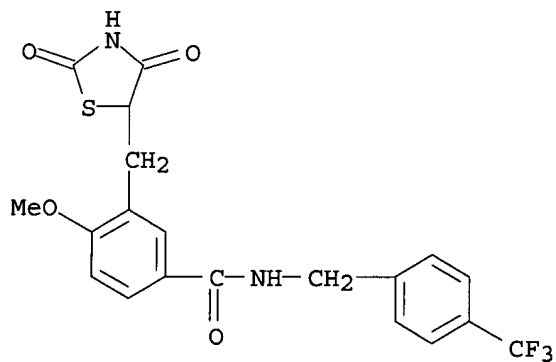
RN 185808-51-9 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-N-[(4-fluorophenyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)

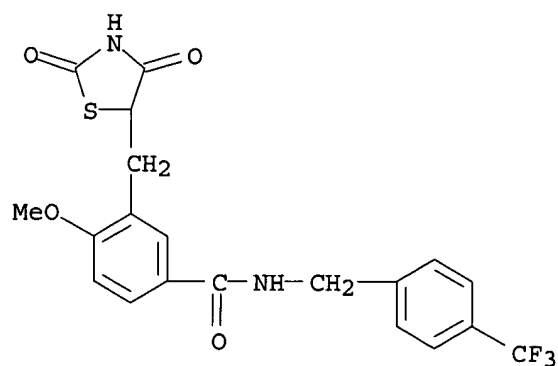


RN 185808-52-0 CAPLUS

CN Benzamide, 3-[(2,4-dioxo-5-thiazolidinyl)methyl]-4-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)

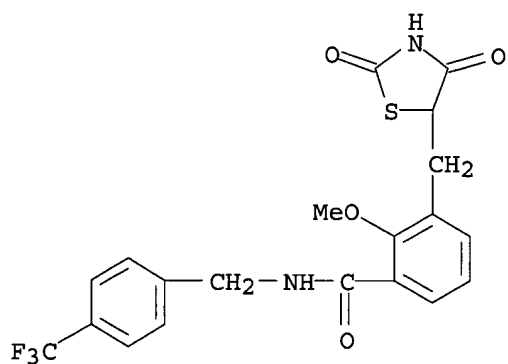


10049937



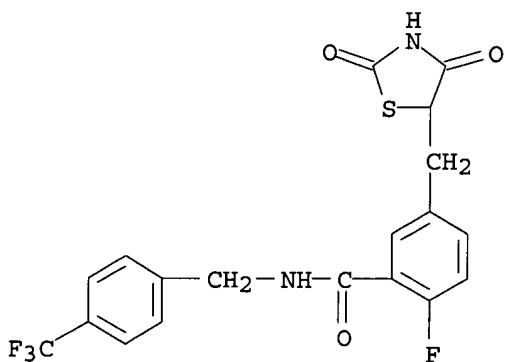
RN 185808-55-3 CAPLUS

CN Benzamide, 3-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



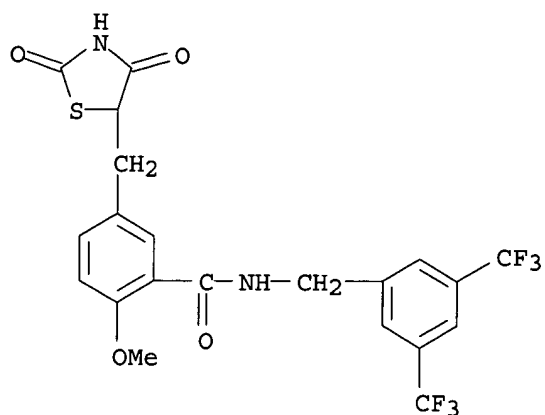
RN 185808-59-7 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-fluoro-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



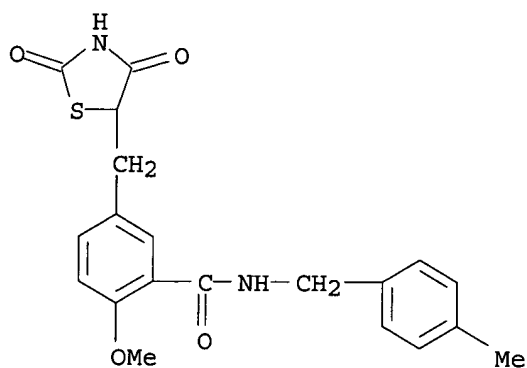
RN 185808-62-2 CAPLUS

CN Benzamide, N-[[3,5-bis(trifluoromethyl)phenyl]methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



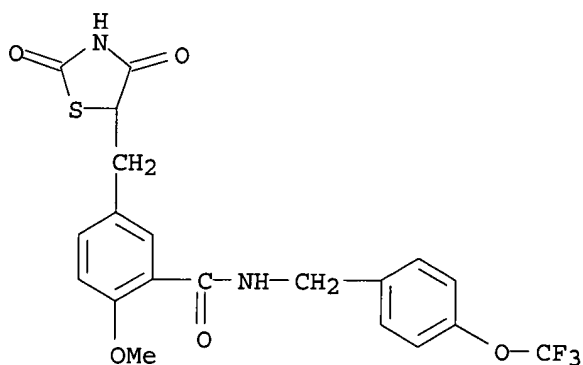
RN 185808-63-3 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-methylphenyl)methyl]- (9CI) (CA INDEX NAME)



RN 185808-64-4 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethoxy)phenyl]methyl]- (9CI) (CA INDEX NAME)

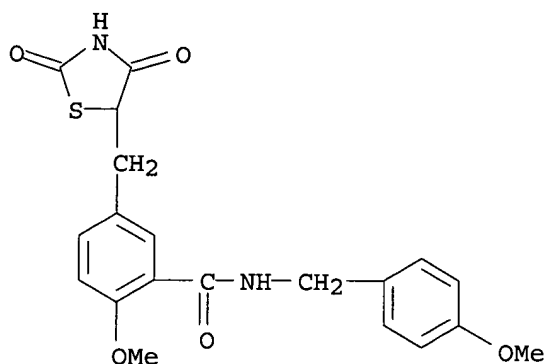


RN 185808-65-5 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-

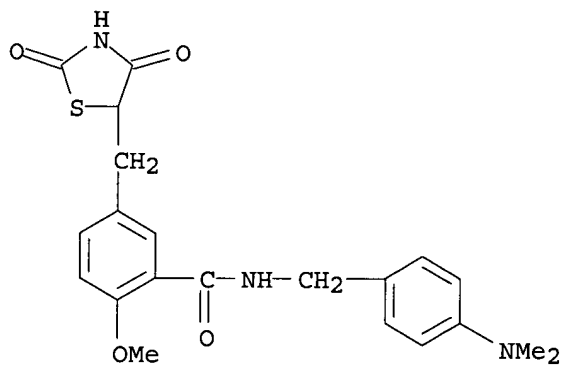
10049937

methoxyphenyl)methyl] - (9CI) (CA INDEX NAME)



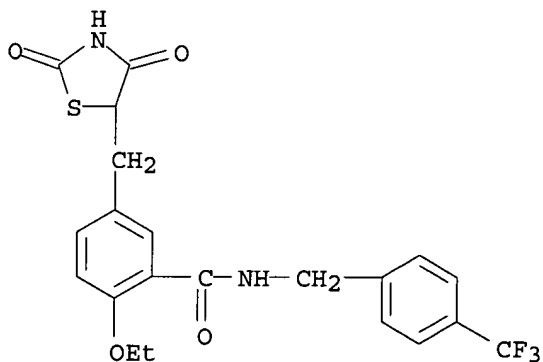
RN 185808-67-7 CAPLUS

CN Benzamide, N-[[4-(dimethylamino)phenyl]methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



RN 185808-68-8 CAPLUS

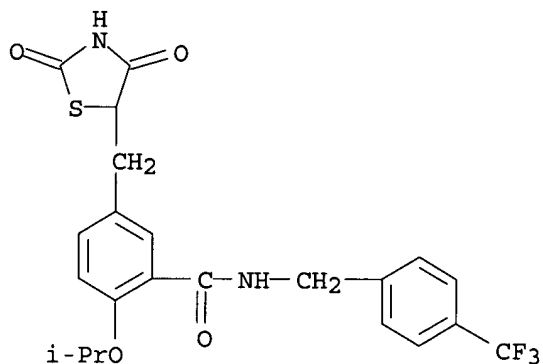
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-ethoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 185808-70-2 CAPLUS

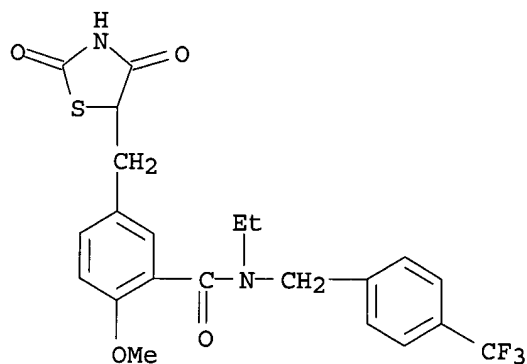
10049937

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-(1-methylethoxy)-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



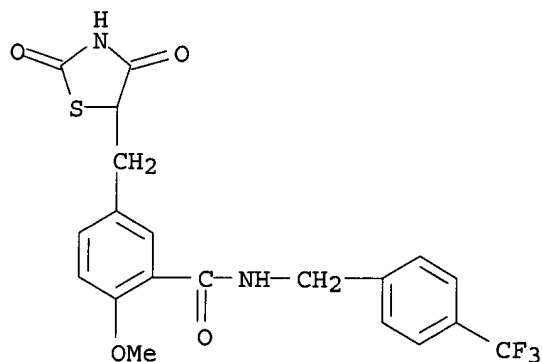
RN 186312-86-7 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-N-ethyl-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 213252-19-8 CAPLUS

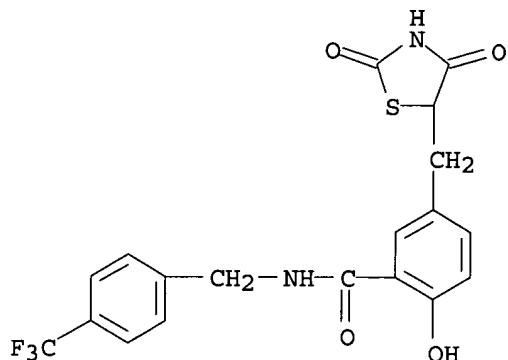
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



10049937

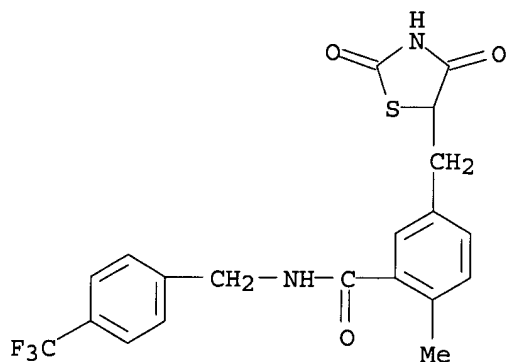
RN 223508-81-4 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-hydroxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 223508-82-5 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methyl-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 38 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1998:784882 CAPLUS

DOCUMENT NUMBER: 130:148506

TITLE: A novel insulin sensitizer acts as a coligand for peroxisome proliferator-activated receptor-.alpha. (PPAR-.alpha.) and PPAR-.gamma.: effect of PPAR-.alpha. activation on abnormal lipid metabolism in liver of Zucker fatty rats

AUTHOR(S): Murakami, Koji; Tobe, Kazuyuki; Ide, Tomohiro; Mochizuki, Toshiro; Ohashi, Mitsuo; Akanuma, Yasuo; Yazaki, Yoshio; Kadowaki, Takashi

CORPORATE SOURCE: Third Department of Internal Medicine, Faculty of Medicine, University of Tokyo, Tokyo, 113, Japan

SOURCE: Diabetes (1998), 47(12), 1841-1847

CODEN: DIAEAZ; ISSN: 0012-1797

PUBLISHER: American Diabetes Association

DOCUMENT TYPE: Journal
 LANGUAGE: English

AB We investigated the biol. activity of a novel thiazolidinedione (TZD) deriv., KRP-297, and the mol. basis of this activity. When administered to obese Zucker fatty rats (obese rats) at 10 mg/kg for 2 wk, KRP-297, unlike BRL-49653, restored reduced lipid oxidn., i.e., CO₂ and ketone body prodn. from [14C]palmitic acid, in the liver by 39% (P < 0.05) and 57% (P < 0.01), resp. KRP-297 was also significantly more effective than BRL-49653 in the inhibition of enhanced lipogenesis and triglyceride accumulation in the liver. To understand the mol. basis of the biol. effects of KRP-297, we examd. the effect on peroxisome proliferator-activated receptor (PPAR) isoforms, which may play key roles in lipid metab. Unlike classical TZD derivs., KRP-297 activated both PPAR-.alpha. and PPAR-.gamma., with median effective concns. of 1.0 and 0.8 .mu.mol/L, resp. Moreover, radiolabeled [3H]KRP-297 bound directly to PPAR-.alpha. and PPAR-.gamma. with dissocn. consts. of 228 and 326 nmol/L, resp. Concomitantly, KRP-297, but not BRL-49653, increased the mRNA and the activity (1.5-fold [P < 0.01] and 1.8-fold [P < 0.05], resp.) of acyl-CoA oxidase, which has been reported to be regulated by PPAR-.alpha., in the liver. By contrast, KRP-297 (P < 0.05) was less potent than BRL-49653 (P < 0.01) in inducing the PPAR-.gamma.-regulated ap2 gene mRNA expression in the adipose tissues. These results suggest that PPAR-.alpha. agonism has a protective effect against abnormal lipid metab. in liver of obese rats.

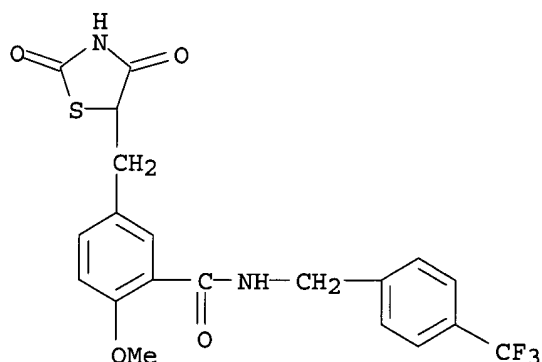
IT 213252-19-8, KRP 297

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(effect of PPAR-.alpha. activation by insulin sensitizer, thiazolidinedione deriv. KRP-297, on abnormal lipid metab. in liver of Zucker fatty rats)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 35 THERE ARE 35 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 39 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1998:421607 CAPLUS

DOCUMENT NUMBER: 129:239719

TITLE: Effects of PPAR.alpha. activation on liver lipid metabolism in Zucker fatty rats

AUTHOR(S): Ide, Tomohiro; Murakami, Koji; Tobe, Kazuyuki; Mochizuki, Toshiro; Ohashi, Mitsuo; Akanuma, Yasuo; Kadowaki, Takashi; Yazaki, Yoshio

CORPORATE SOURCE: Cent. Res. Lab., Kyorin Pharm. Co., Ltd., Tochigi, 329-01, Japan

SOURCE: Diabetes Frontier (1998), 9(3), 345-346
CODEN: DIFREZ; ISSN: 0915-6593

PUBLISHER: Medikaru Rebyusha

DOCUMENT TYPE: Journal

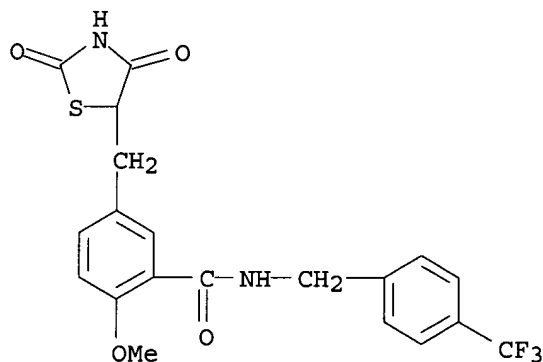
LANGUAGE: Japanese

AB Oral administration of KRP-297 or BRL-49653 with high affinity to PPAR .alpha. to Zucker fatty (obese) rats and to control lean rats for 2 wk significantly lowered the blood glucose, insulin, triglyceride, and free fatty acid levels in the obese rats. KRP-297 and BRL-49653 also suppressed the increase in triglyceride accumulation and fatty acid biosynthesis activity in the liver of the obese rats as compared to the lean rats. In contrast, the markedly reduced activity of the hepatic acyl-CoA oxidase in the obese rats was markedly recovered by the administration. The results suggest that the activation of PPAR .alpha. by KRP-297 or BRL-49653 (ligand) might have inhibitory action on the hepatic triglyceride accumulation and lipid metab. abnormality in the obese rats.

IT 213252-19-8, KRP 297
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)
(effects of PPAR.alpha. activation on liver lipid metab. in Zucker fatty rats)

RN 213252-19-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



L4 ANSWER 40 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1997:116453 CAPLUS

DOCUMENT NUMBER: 126:157499

TITLE: Preparation of N-substituted dioxothiazolidylbenzamide derivatives as blood sugar lowering agents

INVENTOR(S): Maeda, Toshio; Nomura, Masahiro; Awano, Katsuya; Kinoshita, Susumu; Sato, Hiroya; Murakami, Koji; Tsunoda, Masaki

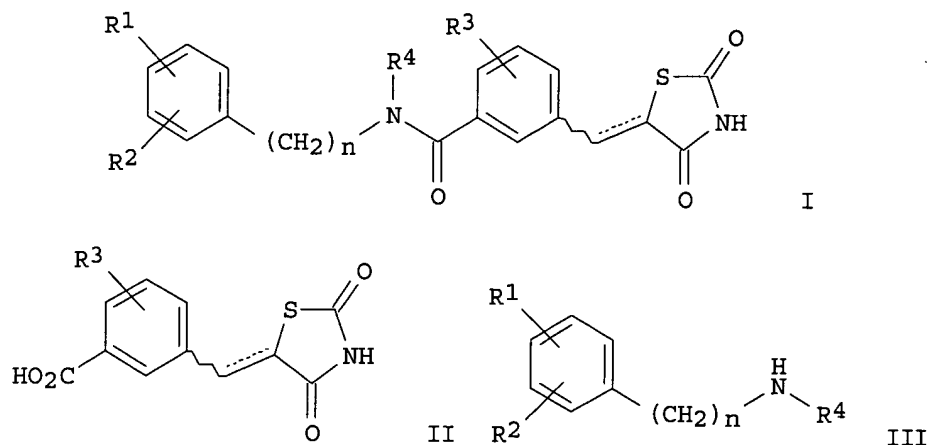
PATENT ASSIGNEE(S): Kyorin Seiyaku Kk, Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 11 pp.
CODEN: JKXXAF

DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 08333355	A2	19961217	JP 1995-159782	19950602
PRIORITY APPLN. INFO.:			JP 1995-159782	19950602
OTHER SOURCE(S):	MARPAT 126:157499			

GI



AB The title compds. (I; R1, R2 = H, C1-4 alkyl, C1-3 alkoxy, haloalkoxy, or haloalkyl, halo, OH, NO2, etc.; R3 = H, C1-3 alkoxy, halo, OH; R4 = H, C1-4 alkyl; dotted line = single or double bond; n = 0-2) are prepd. by reacting benzoic acid derivs. (II; R3, dotted line = same as above) with amines (III; R1, R2, R4, n = same as above). I, possessing blood sugar and lipid lowering activities, are useful for diabetes mellitus and hyperlipemia. Thus, 5-(2,4-dioxothiazolidin-5-ylidene)methyl-2-methoxybenzoic acid was reacted with 4-tert-butylaniline in the presence of Et3N and NCP(O)(OEt)2 to give 99% I (R1 = 4-tert-BuC6H4, R3 = 2-MeO, R2 = R4 = H, dotted line = double bond, n = 0). I (R1 = R2 = 4-CF3, R3 = 6-MeO, R4 = Et, dotted line = single bond, n = 1) at 10 mg/kg showed 31% blood sugar lowering activity when tested on mouses p.o. in vivo.

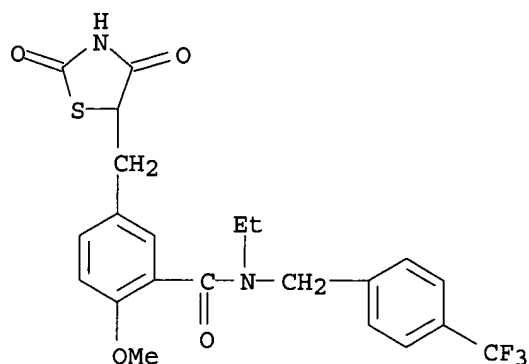
IT 186312-86-7P 186312-87-8P 186312-89-0P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (prepn. of N-substituted dioxothiazolidinylbenzamide derivs. as blood sugar lowering agents)

RN 186312-86-7 CAPLUS

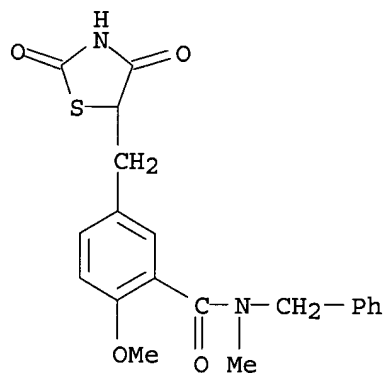
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-N-ethyl-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)

10049937



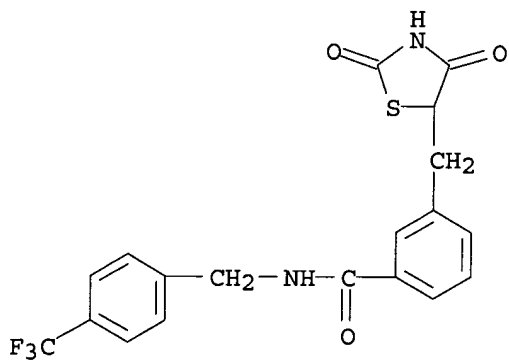
RN 186312-87-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-methyl-N-(phenylmethyl)- (9CI) (CA INDEX NAME)



RN 186312-89-0 CAPLUS

CN Benzamide, 3-[(2,4-dioxo-5-thiazolidinyl)methyl]-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



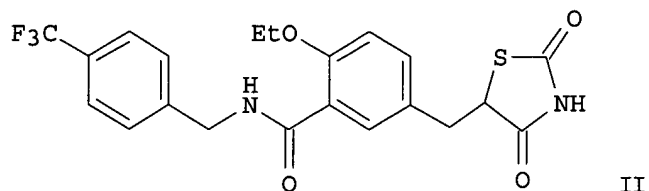
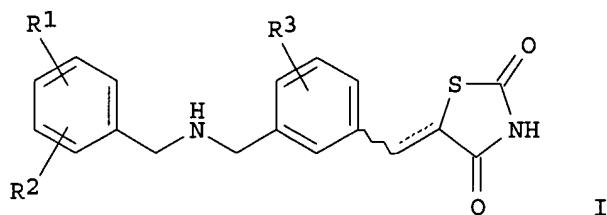
L4 ANSWER 41 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1997:85180 CAPLUS

DOCUMENT NUMBER: 126:104076

TITLE: Preparation of N-benzylthiazolidylbenzamide derivatives as antidiabetics and hypolipemics
 INVENTOR(S): Maeda, Toshio; Nomura, Masahiro; Awano, Katsuya; Kinoshita, Susumu; Satoh, Hiroya; Murakami, Koji; Tsunoda, Masaki
 PATENT ASSIGNEE(S): Kyorin Pharmaceutical Co., Ltd., Japan; Maeda, Toshio; Nomura, Masahiro; Awano, Katsuya; Kinoshita, Susumu; Satoh, Hiroya; Murakami, Koji; Tsunoda, Masaki
 SOURCE: PCT Int. Appl., 40 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9638428	A1	19961205	WO 1996-JP1459	19960530
W: AU, CA, CN, HU, KR, US				
RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
JP 09048771	A2	19970218	JP 1996-153139	19960524
JP 3144624	B2	20010312		
JP 2001139565	A2	20010522	JP 2000-350367	19960524
CA 2220698	AA	19961205	CA 1996-2220698	19960530
AU 9658446	A1	19961218	AU 1996-58446	19960530
AU 698896	B2	19981112		
EP 846693	A1	19980610	EP 1996-920002	19960530
EP 846693	B1	20020123		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, PT, IE, FI				
CN 1186489	A	19980701	CN 1996-194390	19960530
CN 1069901	B	20010822		
AT 212341	E	20020215	AT 1996-920002	19960530
ES 2170858	T3	20020816	ES 1996-920002	19960530
TW 400328	B	20000801	TW 1996-85106555	19960601
US 6030990	A	20000229	US 1997-952672	19971202
US 6001862	A	19991214	US 1999-292955	19990416
US 6147101	A	20001114	US 2000-482268	20000113
CN 1336366	A	20020220	CN 2000-130138	20001017
PRIORITY APPLN. INFO.:			JP 1995-159781	A 19950602
			JP 1996-153139	A 19960524
			WO 1996-JP1459	W 19960530
OTHER SOURCE(S):		MARPAT 126:104076		
GI				



AB Novel N-benzylthiazolidine-2,4-dione derivs. represented by general formula I [R1 and R2 are the same or different and each represents hydrogen, lower (C1-4) alkyl, lower (C1-3) alkoxy, lower (C1-3) haloalkyl, lower (C1-3) haloalkoxy, halogeno, hydroxy, nitro, amino optionally substituted by lower (C1-3) alkyl or a heterocycle, or R1 and R2 may be bonded to each other to form methylenedioxy; R3 represents lower (C1-3) alkoxy, hydroxy or halogeno; and the dotted line represents a double or single bond] are prepd. The title compd. II at 10 mg/kg gave 37% decrease in blood sugar in obese mice.

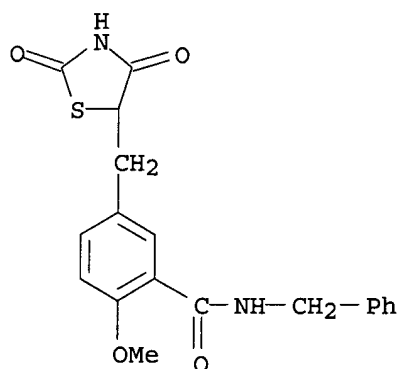
IT 185808-38-2P 185808-40-6P 185808-42-8P
 185808-45-1P 185808-49-5P 185808-51-9P
 185808-52-0P 185808-54-2P 185808-55-3P
 185808-59-7P 185808-62-2P 185808-63-3P
 185808-64-4P 185808-65-5P 185808-67-7P
 185808-68-8P 185808-70-2P 185808-71-3P
 185808-72-4P 185808-73-5P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of N-benzylthiazolidine-2,4-dione derivs. as antidiabetics and hypolipemics)

RN 185808-38-2 CAPLUS

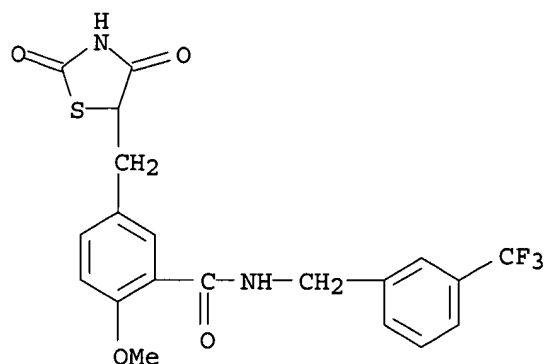
CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-(phenylmethyl)- (9CI) (CA INDEX NAME)



10049937

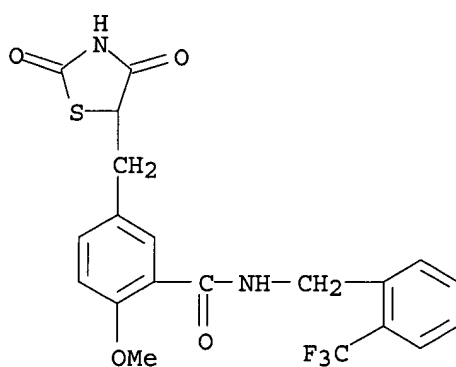
RN 185808-40-6 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[3-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



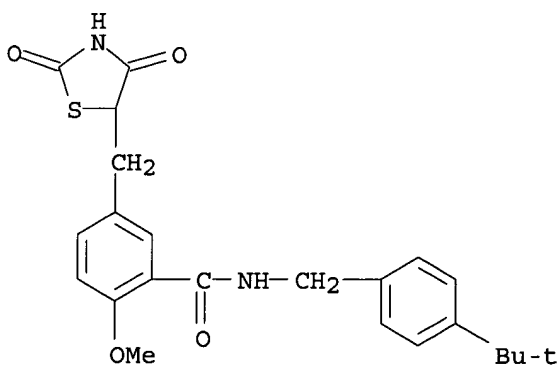
RN 185808-42-8 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[2-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 185808-45-1 CAPLUS

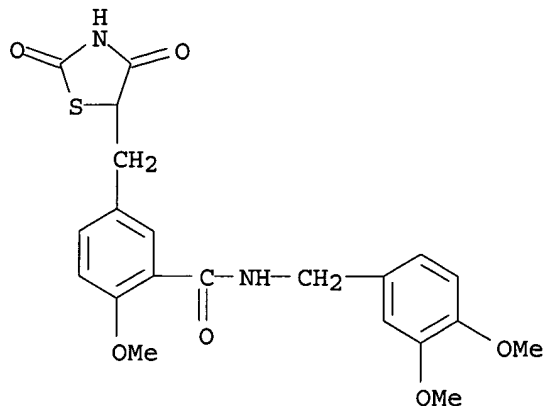
CN Benzamide, N-[[4-(1,1-dimethylethyl)phenyl]methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



10049937

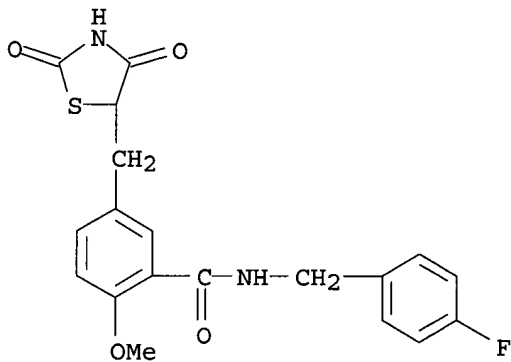
RN 185808-49-5 CAPLUS

CN Benzamide, N-[(3,4-dimethoxyphenyl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



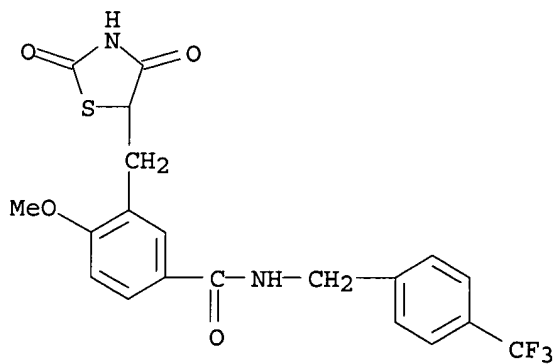
RN 185808-51-9 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-N-[(4-fluorophenyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)

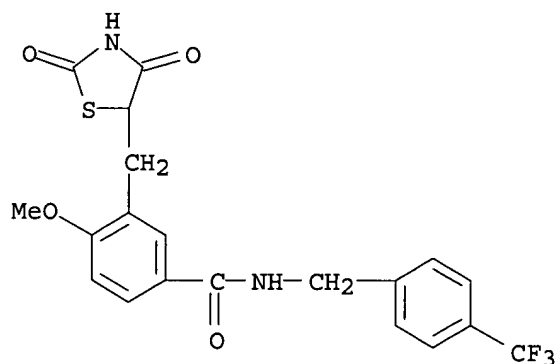


RN 185808-52-0 CAPLUS

CN Benzamide, 3-[(2,4-dioxo-5-thiazolidinyl)methyl]-4-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)

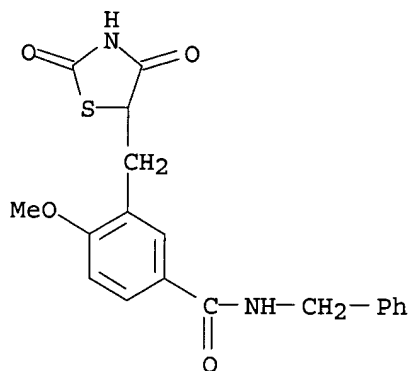


10049937



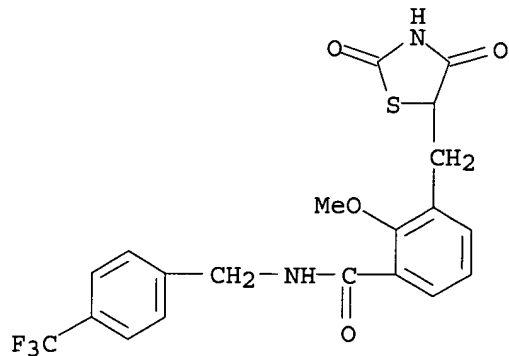
RN 185808-54-2 CAPLUS

CN Benzamide, 3-[(2,4-dioxo-5-thiazolidinyl)methyl]-4-methoxy-N-(phenylmethyl)- (9CI) (CA INDEX NAME)



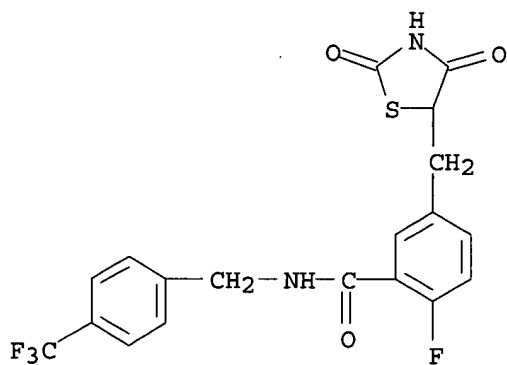
RN 185808-55-3 CAPLUS

CN Benzamide, 3-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



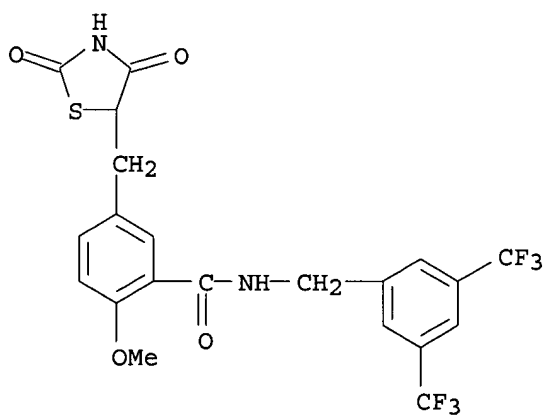
RN 185808-59-7 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-fluoro-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



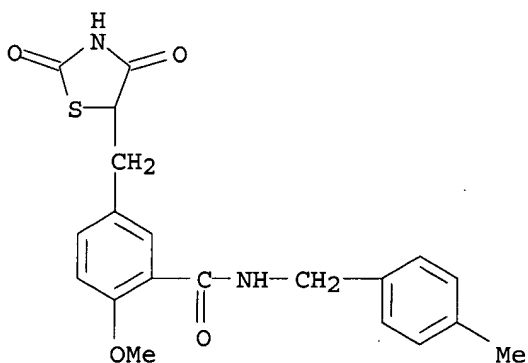
RN 185808-62-2 CAPLUS

CN Benzamide, N-[[3,5-bis(trifluoromethyl)phenyl]methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



RN 185808-63-3 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-methylphenyl)methyl]- (9CI) (CA INDEX NAME)

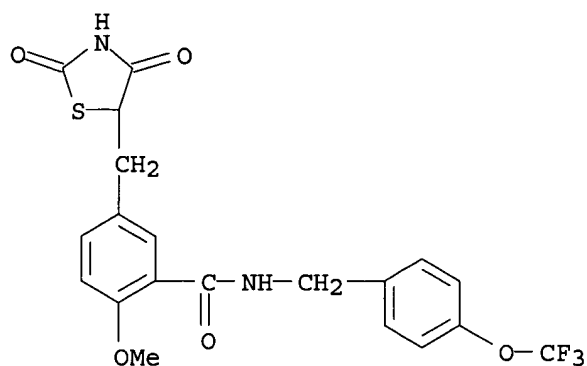


RN 185808-64-4 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-

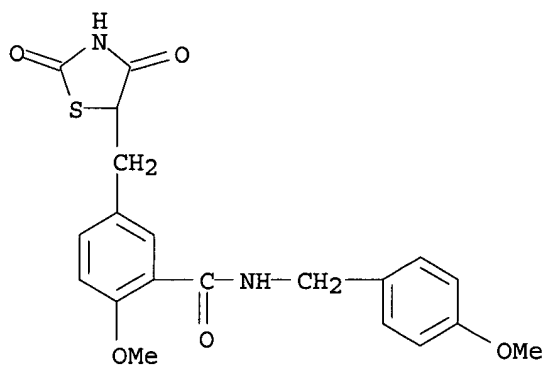
10049937

(trifluoromethoxy)phenyl)methyl] - (9CI) (CA INDEX NAME)



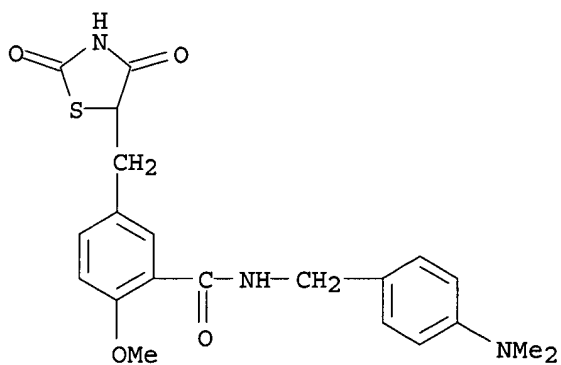
RN 185808-65-5 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[(4-methoxyphenyl)methyl] - (9CI) (CA INDEX NAME)



RN 185808-67-7 CAPLUS

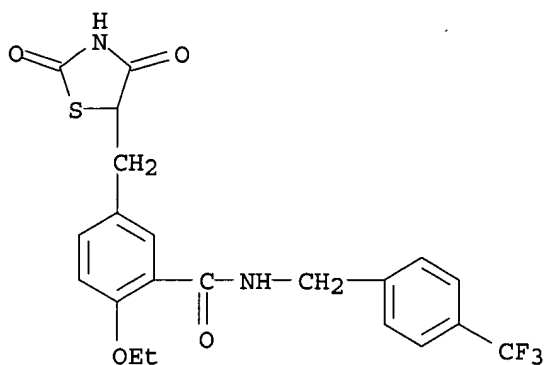
CN Benzamide, N-[[4-(dimethylamino)phenyl)methyl]-5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy- (9CI) (CA INDEX NAME)



RN 185808-68-8 CAPLUS

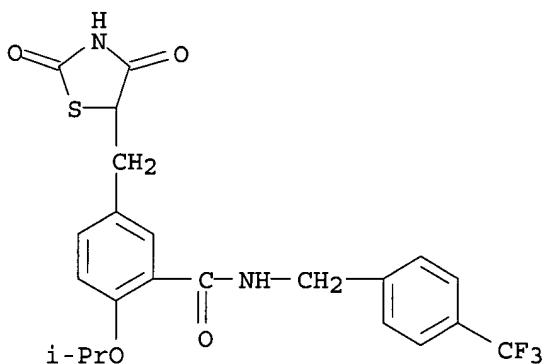
10049937

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-ethoxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



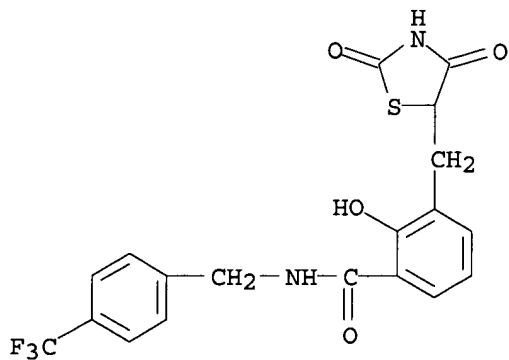
RN 185808-70-2 CAPLUS

CN Benzamide, 5-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-(1-methylethoxy)-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



RN 185808-71-3 CAPLUS

CN Benzamide, 3-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-hydroxy-N-[[4-(trifluoromethyl)phenyl]methyl]- (9CI) (CA INDEX NAME)



10049937

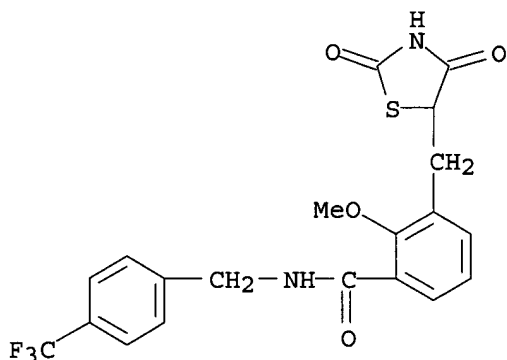
RN 185808-72-4 CAPLUS

CN Benzamide, 3-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]-, compd. with (S)-.alpha.-methylbenzenemethanamine (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 185808-55-3

CMF C20 H17 F3 N2 O4 S

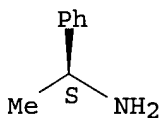


CM 2

CRN 2627-86-3

CMF C8 H11 N

Absolute stereochemistry.



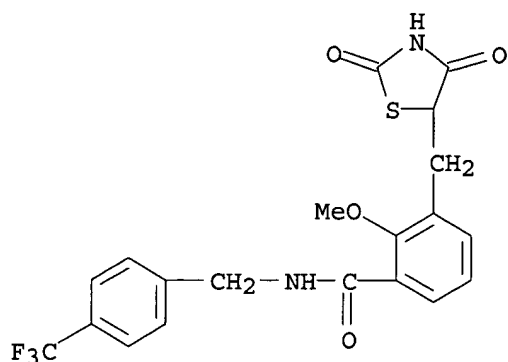
RN 185808-73-5 CAPLUS

CN Benzamide, 3-[(2,4-dioxo-5-thiazolidinyl)methyl]-2-methoxy-N-[[4-(trifluoromethyl)phenyl]methyl]-, compd. with (R)-.alpha.-methylbenzenemethanamine (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 185808-55-3

CMF C20 H17 F3 N2 O4 S

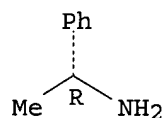


CM 2

CRN 3886-69-9

CMF C8 H11 N

Absolute stereochemistry.



L4 ANSWER 42 OF 42 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 1996:537365 CAPLUS

DOCUMENT NUMBER: 125:195637

TITLE: Preparation of dioxothiazolidine derivatives as hypoglycemics and aldose reductase inhibitors

INVENTOR(S): Matsushima, Hiroaki; Sugizaki, Myoshi; Myaoka, Shozo

PATENT ASSIGNEE(S): Terumo Corp, Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 31 pp.

CODEN: JKXXAF

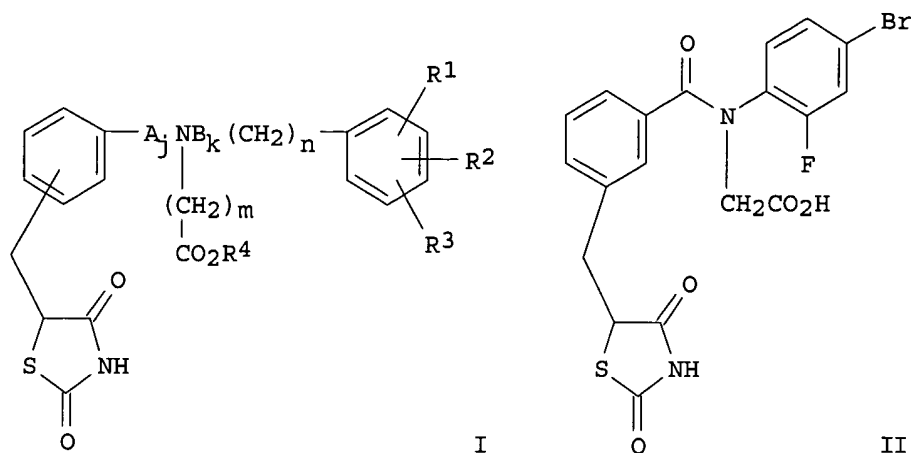
DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
-----	----	-----	-----	-----
JP 08143556	A2	19960604	JP 1994-280957	19941115
PRIORITY APPLN. INFO.:			JP 1994-280957	19941115
OTHER SOURCE(S):	MARPAT 125:195637			
GI				



AB The title compds. I [A, B = carbonyl, etc.; R₄ = H, alkyl; R₁ - R₃ = H, halo, etc.; j, k = 0 or 1; m = 1 or 2; n = 0 or 1] are prepd. The title compd. II (prepn. given) in vitro showed IC₅₀ of 8.32 x 10⁻⁷ M against aldose reductase. II also showed hypoglycemic activity.

IT **180631-42-9P**

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(prepn. of dioxothiazolidine derivs. as hypoglycemics and aldose reductase inhibitors)

RN 180631-42-9 CAPLUS

CN Glycine, N-[(4-bromo-2-fluorophenyl)methyl]-N-[3-[(2,4-dioxo-5-thiazolidinyl)methyl]benzoyl]- (9CI) (CA INDEX NAME)

